Refine Search

Search Results -

Term	Documents
HAEMOPHILIA	535
HAEMOPHILIUM	2
HAEMOPHILIUMS	0
HAEMOPHILIAS	13
(HAEMOPHILIA AND 18).PGPB,USPT.	43
(L18 AND HAEMOPHILIA).PGPB,USPT.	43

US Pre-Grant Publication Full-Text Database US Patents Full-Text Database

Database:

US OCR Full-Text Database EPO Abstracts Database JPO Abstracts Database Derwent World Patents Index IBM Technical Disclosure Bulletins

Search:

L19		The second secon	Refine Search
Recall Text 👄	Clear		Interrupt

Search History

DATE: Thursday, October 28, 2004 Printable Copy Create Case

Set Name side by side		Hit Count Set Name result set				
DB=PC	GPB,USPT; THES=ASSIGNEE; PLUR=YES;	OP=ADJ				
<u>L19</u>	L18 and haemophilia	43	<u>L19</u>			
L18	L17 and composition	171	<u>L18</u>			
<u>L17</u>	FIX and FVIII	220	<u>L17</u>			
<u>L16</u>	L15and treatment	0	<u>L16</u>			
<u>L15</u>	FIX and haemophilia	80	<u>L15</u>			
<u>L14</u>	factor IX and treatment of haemophilia	0	<u>L14</u>			
<u>L13</u>	factor IX same treatement of haemephilia	0	<u>L13</u>			
<u>L12</u>	L10 and treatment	118	<u>L12</u>			
<u>L11</u>	L10 and treatement	. 0	<u>L11</u>			

<u>L9</u>	factor IX composition and phospholipid	7	<u>L9</u>
<u>L8</u>	L7 and py<2003	60	<u>L8</u>
<u>L7</u>	L6 and pharmaceutical composition	60	<u>L7</u>
<u>L6</u>	L4 and haemophilia and phospholipid	69	<u>L6</u>
<u>L5</u>	L4 and haemophilia	69	<u>L5</u>
<u>L4</u>	L3 and factor VIII	612	<u>L4</u>
<u>L3</u>	L2 and pharmaceutical	1076	<u>L3</u>
<u>L2</u>	L1 and composition	1240	<u>L2</u>
<u>L1</u>	factor IX and phospholipid	1336	<u>L1</u>

END OF SEARCH HISTORY

<u>L10</u>	L9 and haemophilia	124	<u>L10</u>
<u>L9</u>	factor IX same factor VIII	1921	<u>L9</u>
<u>L8</u>	composition of factor IX and factor VIII	0	<u>L8</u>
<u>L7</u>	L2 and haemophilia	6	<u>L7</u>
<u>L6</u>	L5 and haemophilia	17	<u>L6</u>
<u>L5</u>	(factor IX and factor VIII)same composition	194	<u>L5</u>
<u>L4</u>	L3 and py<2004	6	<u>L4</u>
<u>L3</u>	L2 and haemophilia	6	<u>L3</u>
<u>L2</u>	composition same factor IXa and factor VIII	53	<u>L2</u>
<u>L1</u>	composition FIXa and FVIII	0	<u>L1</u>

END OF SEARCH HISTORY

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 NEWS 3 Jul 12 BEILSTEIN enhanced with new display and select options,
                  resulting in a closer connection to BABS
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      4 AUG 02 IFIPAT/IFIUDB/IFICDB reloaded with new search and display
                  fields
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      7
         AUG 27
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         AUG 27
                  status data from INPADOC
         SEP 01
                 INPADOC: New family current-awareness alert (SDI) available
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                 New pricing for the Save Answers for SciFinder Wizard within
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        SEP 01
                  STN Express with Discover!
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         SEP 01
NEWS 11
         SEP 27
NEWS 12
                 STANDARDS will no longer be available on STN
         SEP 27 SWETSCAN will no longer be available on STN
NEWS 13
NEWS EXPRESS
              JULY 30 CURRENT WINDOWS VERSION IS V7.01, CURRENT
               MACINTOSH VERSION IS V6.0c(ENG) AND V6.0Jc(JP),
               AND CURRENT DISCOVER FILE IS DATED 11 AUGUST 2004
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               STN Operating Hours Plus Help Desk Availability
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               General Internet Information
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               Welcome Banner and News Items
NEWS PHONE
              Direct Dial and Telecommunication Network Access to STN
NEWS WWW
               CAS World Wide Web Site (general information)
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Enter x:

Enter x:

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=> index bioscience
FILE 'DRUGMONOG' ACCESS NOT AUTHORIZED
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El

SINCE FILE TOTAL ENTRY SESSION 0.21 0.21

3.06

FULL ESTIMATED COST

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 11:02:32 ON 28 OCT 2004

75 FILES IN THE FILE LIST IN STNINDEX

Enter SET DETAIL ON to see search term postings or to view search error messages that display as 0* with SET DETAIL OFF.

- => s composit? and factor IX and facror VIII
 - 17 FILES SEARCHED...
 - 27 FILES SEARCHED...
 - 46 FILES SEARCHED...
 - 61 FILES SEARCHED...
 - 1 FILE USPATFULL
 - 74 FILES SEARCHED...
 - 1 FILES HAVE ONE OR MORE ANSWERS, 75 FILES SEARCHED IN STNINDEX
- L1 QUE COMPOSIT? AND FACTOR IX AND FACROR VIII
- => d rank
- F1 1 USPATFULL

=> file F1

COST IN U.S. DOLLARS SINCE FILE TOTAL ENTRY SESSION

FULL ESTIMATED COST 2.85

FILE 'USPATFULL' ENTERED AT 11:05:46 ON 28 OCT 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE COVERS 1971 TO PATENT PUBLICATION DATE: 26 Oct 2004 (20041026/PD) FILE LAST UPDATED: 26 Oct 2004 (20041026/ED) HIGHEST GRANTED PATENT NUMBER: US6810528

HIGHEST APPLICATION PUBLICATION NUMBER: US2004210975 CA INDEXING IS CURRENT THROUGH 26 Oct 2004 (20041026/UPCA)

ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 26 Oct 2004 (20041026/PD)

REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2004

USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2004

- >>> USPAT2 is now available. USPATFULL contains full text of the <<<
- >>> original, i.e., the earliest published granted patents or
- >>> applications. USPAT2 contains full text of the latest US <<<
- >>> publications, starting in 2001, for the inventions covered in <<<
- >>> USPATFULL. A USPATFULL record contains not only the original
- >>> published document but also a list of any subsequent
- >>> publications. The publication number, patent kind code, and <<<
- >>> publication date for all the US publications for an invention <<<
- >>> are displayed in the PI (Patent Information) field of USPATFULL <<<
- >>> records and may be searched in standard search fields, e.g., /PN, <<< >>> /PK, etc.
- >>> USPATFULL and USPAT2 can be accessed and searched together

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>>> through the new cluster USPATALL. Type FILE USPATALL to
>>> enter this cluster.
                                                                         <<<
>>> Use USPATALL when searching terms such as patent assignees,
    classifications, or claims, that may potentially change from
>>> the earliest to the latest publication.
This file contains CAS Registry Numbers for easy and accurate
substance identification.
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        483165 FACTORS
        788251 FACTOR
                 (FACTOR OR FACTORS)
        112561 IX
             1 IXES
        112562 IX
                  (IX OR IXES)
          3425 FACTOR IX
                  (FACTOR(W)IX)
             4 FACROR
        143803 VIII
            25 VIIIS
        143806 VIII
                  (VIII OR VIIIS)
             1 FACROR VIII
                  (FACROR(W)VIII)
             1 COMPOSIT? AND FACTOR IX AND FACROR VIII
L2
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L1 HAS NO ANSWERS
                QUE ABB=ON PLU=ON COMPOSIT? AND FACTOR IX AND FACROR V
L1
                III
=> d L2 1 bib, abs
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       2004:132900 USPATFULL
AN
       Methods for sterilizing biological materials
TΙ
       MacPhee, Martin J., Montgomery Village, MD, UNITED STATES
ΤN
       Kent, Randall S., Thousand Oaks, CA, UNITED STATES Horton, Edward A., Toronto, CANADA
       Beall, Dawson, Gaithersburg, MD, UNITED STATES
       Clearant, Inc. (U.S. corporation)
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PΙ
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                                20040527
                                20031029 (10)
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       Continuation-in-part of Ser. No. US 1995-573149, filed on 15 Dec 1995,
       GRANTED, Pat. No. US 6171549 Continuation-in-part of Ser. No. WO
       1994-CA401, filed on 22 Jul 1994, UNKNOWN Continuation-in-part of Ser.
       No. US 1993-95698, filed on 22 Jul 1993, GRANTED, Pat. No. US 5362442
DT
       Utility
       APPLICATION
FS
       FLESHNER & KIM, LLP, P.O. BOX 221200, CHANTILLY, VA, 20153
LREP
CLMN
       Number of Claims: 36
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 1095
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Methods are disclosed for sterilizing biological products to reduce the
       level of active biological contaminants such as viruses, bacteria,
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yeasts, molds, mycoplasmas and parasites.
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CAS INDEXING IS AVAILABLE FOR THIS PATENT.

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(FILE 'HOME' ENTERED AT 11:02:09 ON 28 OCT 2004)
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INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 11:02:32 ON 28 OCT 2004 SEA COMPOSIT? AND FACTOR IX AND FACROR VIII

1 FILE USPATFULL

L1 QUE COMPOSIT? AND FACTOR IX AND FACROR VIII

FILE 'USPATFULL' ENTERED AT 11:05:46 ON 28 OCT 2004

L2 1 S L1

=> s L1 and haemophilia

1009635 COMPOSIT?

509873 FACTOR

483165 FACTORS

788251 FACTOR

(FACTOR OR FACTORS)

112561 IX

1 IXES

112562 IX

(IX OR IXES)

3425 FACTOR IX

(FACTOR(W)IX)

4 FACROR

143803 VIII

25 VIIIS

143806 VIII

(VIII OR VIIIS)

1 FACROR VIII

(FACROR(W)VIII)

458 HAEMOPHILIA

13 HAEMOPHILIAS

468 HAEMOPHILIA

(HAEMOPHILIA OR HAEMOPHILIAS)

L3 0 L1 AND HAEMOPHILIA

=> s Composition and FIX and FVIII

760744 COMPOSITION

465872 COMPOSITIONS

818786 COMPOSITION

(COMPOSITION OR COMPOSITIONS)

131131 FIX

36375 FIXES

155796 FIX

(FIX OR FIXES)

516 FVIII

1 FVIIIS

516 FVIII

(FVIII OR FVIIIS)

L4 171 COMPOSITION AND FIX AND FVIII

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F1 1 USPATFULL

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=> d L4 1 bib, abs
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       2004:267309 USPATFULL
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ΤТ
       Mini-Ad vector for immunization
ΤN
       Zhang, Wei-Wei, Libertyville, IL, UNITED STATES
       Alemany, Ramon, Grayslake, IL, UNITED STATES
       Dai, Yifan, Grayslake, IL, UNITED STATES
       Josephs, Steven, Grayslake, IL, UNITED STATES
       Balaque, Cristina, Grayslake, IL, UNITED STATES
       Ayares, David, Blacksburgh, VA, UNITED STATES
       Schneiderman, Richard, Highland Park, IL, UNITED STATES
PΙ
       US 2004208846
                               20041021
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ΑI
       US 2004-837079
                               20040615 (9)
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RLI
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       Jan 1997, ABANDONED
       US 2000-197734P
                          20000418 (60)
PRAI
       US 2000-198501P
                           20000418 (60)
DT
       Utility
FS
       APPLICATION
       MCDONNELL BOEHNEN HULBERT & BERGHOFF LLP, 300 S. WACKER DRIVE, 32ND
LREP
       FLOOR, CHICAGO, IL, 60606
       Number of Claims: 29
CLMN
       Exemplary Claim: 1
ECL
       57 Drawing Page(s)
DRWN
LN.CNT 4461
       The present invention provides a method for treating a disorder such as
AB
       hemophilia. A method of treating hemophilia in a mammal by administering
       recombinant virus virions comprising a nucleotide sequence having an
       adenoviral inverted terminal repeat fusion sequence, a packaging signal,
       a transcriptional control region, and a nucleic acid encoding a
       therapeutic protein such as FVIII. In addition, the DNA
       molecule does not encode an adenoviral protein. It is preferred that the
       virions be administered to the mammal under conditions that result in
       the expression of the therapeutic protein at a level that provides a
       therapeutic effect in said mammal.
=> s FVIII and treatement and haemophilia
           516 FVIII
             1 FVIIIS
           516 FVIII
                 (FVIII OR FVIIIS)
          1002 TREATEMENT
            53 TREATEMENTS
          1052 TREATEMENT
                 (TREATEMENT OR TREATEMENTS)
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(FVIII OR FVIIIS)

1002 TREATEMENT
53 TREATEMENTS
1052 TREATEMENT OR TREATEMENTS)

458 HAEMOPHILIA
13 HAEMOPHILIAS
468 HAEMOPHILIA OR HAEMOPHILIAS)
L5 0 FVIII AND TREATEMENT AND HAEMOPHILIA
=> sfile caplus

=> stile captus
SFILE IS NOT A RECOGNIZED COMMAND
The previous command name entered was not recognized by the system.
For a list of commands available to you in the current file, enter
"HELP COMMANDS" at an arrow prompt (=>).

COST IN U.S. DOLLARS

SINCE FILE

TOTAL SESSION

FULL ESTIMATED COST

ENTRY 17.18

20.24

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FILE COVERS 1907 - 28 Oct 2004 VOL 141 ISS 18 FILE LAST UPDATED: 27 Oct 2004 (20041027/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s factor FVIII and FIX

859700 FACTOR

762249 FACTORS

1358572 FACTOR

(FACTOR OR FACTORS)

1220 FVIII

1 FVIIIS

1220 FVIII

(FVIII OR FVIIIS)

80 FACTOR FVIII

(FACTOR(W)FVIII)

11554 FIX

2307 FIXES

13733 FIX

(FIX OR FIXES)

 $_{
m L6}$

3 FACTOR FVIII AND FIX

=> d rank

F1

1 USPATFULL

=> file 1

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

FULL ESTIMATED COST

ENTRY 6.34 SESSION 26.58

FILE '1MOBILITY' ENTERED AT 11:12:41 ON 28 OCT 2004 COPYRIGHT (C) 2004 Society of Automotive Engineers, Inc.

FILE COVERS 1906 TO 1 Oct 2004 (20041001/ED)

1MOBILITY and 2MOBILITY, which together comprise the Global Mobility Database, can be accessed and searched together through the file cluster MOBILITY. Type FILE MOBILITY to enter this cluster.

=> file f1 COST IN U.S. DOLLARS

SINCE FILE

TOTAL

FILE 'USPATFULL' ENTERED AT 11:12:54 ON 28 OCT 2004
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FILE COVERS 1971 TO PATENT PUBLICATION DATE: 26 Oct 2004 (20041026/PD)
FILE LAST UPDATED: 26 Oct 2004 (20041026/ED)
HIGHEST GRANTED PATENT NUMBER: US6810528
HIGHEST APPLICATION PUBLICATION NUMBER: US2004210975
CA INDEXING IS CURRENT THROUGH 26 Oct 2004 (20041026/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 26 Oct 2004 (20041026/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2004
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2004

>>> USPAT2 is now available. USPATFULL contains full text of the <<< >>> original, i.e., the earliest published granted patents or <<< >>> applications. USPAT2 contains full text of the latest US <<< >>> publications, starting in 2001, for the inventions covered in <<< >>> USPATFULL. A USPATFULL record contains not only the original <<< >>> published document but also a list of any subsequent <<< publications. The publication number, patent kind code, and <<< >>> publication date for all the US publications for an invention <<< >>> are displayed in the PI (Patent Information) field of USPATFULL <<< >>> records and may be searched in standard search fields, e.g., /PN, <<< >>> /PK, etc. >>> USPATFULL and USPAT2 can be accessed and searched together <<< >>> through the new cluster USPATALL. Type FILE USPATALL to <<< >>> enter this cluster. <<< <<< >>> >>> Use USPATALL when searching terms such as patent assignees, <<< >>> classifications, or claims, that may potentially change from <<< >>> the earliest to the latest publication. <<<

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> s L1 1009635 COMPOSIT? 509873 FACTOR **483165 FACTORS** 788251 FACTOR (FACTOR OR FACTORS) 112561 IX 1 IXES 112562 IX (IX OR IXES) 3425 FACTOR IX (FACTOR (W) IX) 4 FACROR 143803 VIII 25 VIIIS 143806 VIII (VIII OR VIIIS) 1 FACROR VIII (FACROR(W)VIII)

L7 1 COMPOSIT? AND FACTOR IX AND FACROR VIII

=> d his

(FILE 'HOME' ENTERED AT 11:02:09 ON 28 OCT 2004)

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE,

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AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS, BIOTECHAS,
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               SEA COMPOSIT? AND FACTOR IX AND FACROR VIII
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L1
                QUE COMPOSIT? AND FACTOR IX AND FACROR VIII
     FILE 'USPATFULL' ENTERED AT 11:05:46 ON 28 OCT 2004
L2
              1 S L1
              0 S L1 AND HAEMOPHILIA
L_3
L4
            171 S COMPOSITION AND FIX AND FVIII
L5
              O S FVIII AND TREATEMENT AND HAEMOPHILIA
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              3 S FACTOR FVIII AND FIX
L6
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L7
=> $ L7
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            1 IXES
        112562 IX
                 (IX OR IXES)
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        143806 VIII
                 (VIII OR VIIIS)
             1 FACROR VIII
                 (FACROR(W)VIII)
             1 COMPOSIT? AND FACTOR IX AND FACROR VIII
L8
=> d rank
             1 USPATFULL
F1
=> file f1
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COST IN U.S. DOLLARS
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                                                      ENTRY
                                                               SESSION
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                                                                  29.70
FULL ESTIMATED COST
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FILE LAST UPDATED: 26 Oct 2004 (20041026/ED)
HIGHEST GRANTED PATENT NUMBER: US6810528
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ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 26 Oct 2004 (20041026/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2004
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USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2004

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>>> USPAT2 is now available. USPATFULL contains full text of the
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>>> original, i.e., the earliest published granted patents or
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>>> applications. USPAT2 contains full text of the latest US
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     publications, starting in 2001, for the inventions covered in
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>>> USPATFULL. A USPATFULL record contains not only the original
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>>> published document but also a list of any subsequent
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    publications. The publication number, patent kind code, and
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>>> are displayed in the PI (Patent Information) field of USPATFULL
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>>> /PK, etc.
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>>> through the new cluster USPATALL. Type FILE USPATALL to
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>>> enter this cluster.
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>>> Use USPATALL when searching terms such as patent assignees,
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>>> the earliest to the latest publication.
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This file contains CAS Registry Numbers for easy and accurate
substance identification.
=> d L8 1 bib, abs
     ANSWER 1 OF 1 USPATFULL on STN
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AN
       Methods for sterilizing biological materials
TI
       MacPhee, Martin J., Montgomery Village, MD, UNITED STATES
IN
       Kent, Randall S., Thousand Oaks, CA, UNITED STATES Horton, Edward A., Toronto, CANADA
       Beall, Dawson, Gaithersburg, MD, UNITED STATES
       Clearant, Inc. (U.S. corporation)
PΑ
PΙ
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                          A1
                               20040527
                               20031029 (10)
       US 2003-694733
AΙ
                          A 1
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RLI
       Continuation-in-part of Ser. No. US 1995-573149, filed on 15 Dec 1995,
       GRANTED, Pat. No. US 6171549 Continuation-in-part of Ser. No. WO
       1994-CA401, filed on 22 Jul 1994, UNKNOWN Continuation-in-part of Ser.
       No. US 1993-95698, filed on 22 Jul 1993, GRANTED, Pat. No. US 5362442
DТ
       Utility
       APPLICATION
FS
       FLESHNER & KIM, LLP, P.O. BOX 221200, CHANTILLY, VA, 20153
LREP
       Number of Claims: 36
CLMN
       Exemplary Claim: 1
ECL
DRWN
       No Drawings
LN.CNT 1095
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Methods are disclosed for sterilizing biological products to reduce the
       level of active biological contaminants such as viruses, bacteria,
       yeasts, molds, mycoplasmas and parasites.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
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=> s L8 and composition
760744 COMPOSITION
465872 COMPOSITIONS
818786 COMPOSITION
(COMPOSITION OR COMPOSITIONS)
L9 1 L8 AND COMPOSITION
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=> d L9 1 bib,abs
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       Methods for sterilizing biological materials
       MacPhee, Martin J., Montgomery Village, MD, UNITED STATES
TN
       Kent, Randall S., Thousand Oaks, CA, UNITED STATES Horton, Edward A., Toronto, CANADA
       Beall, Dawson, Gaithersburg, MD, UNITED STATES
       Clearant, Inc. (U.S. corporation)
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                          Α1
                                20031029 (10)
       Continuation of Ser. No. US 2000-533547, filed on 23 Mar 2000, PENDING
RLI
       Continuation-in-part of Ser. No. US 1995-573149, filed on 15 Dec 1995,
       GRANTED, Pat. No. US 6171549 Continuation-in-part of Ser. No. WO
       1994-CA401, filed on 22 Jul 1994, UNKNOWN Continuation-in-part of Ser.
       No. US 1993-95698, filed on 22 Jul 1993, GRANTED, Pat. No. US 5362442
DT
       Utility
FS
       APPLICATION
LREP
       FLESHNER & KIM, LLP, P.O. BOX 221200, CHANTILLY, VA, 20153
CLMN
       Number of Claims: 36
ECL
       Exemplary Claim: 1
DRWN
       No Drawings
LN.CNT 1095
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
       Methods are disclosed for sterilizing biological products to reduce the
       level of active biological contaminants such as viruses, bacteria,
       yeasts, molds, mycoplasmas and parasites.
CAS INDEXING IS AVAILABLE FOR THIS PATENT.
=> s L9 and haemophilia
           458 HAEMOPHILIA
            13 HAEMOPHILIAS
           468 HAEMOPHILIA
                  (HAEMOPHILIA OR HAEMOPHILIAS)
L10
             0 L9 AND HAEMOPHILIA
=> s composition of FIX and FVIII
        760744 COMPOSITION
        465872 COMPOSITIONS
        818786 COMPOSITION
                  (COMPOSITION OR COMPOSITIONS)
        131131 FIX
         36375 FIXES
        155796 FIX
                  (FIX OR FIXES)
            85 COMPOSITION OF FIX
                  (COMPOSITION (1W) FIX)
           516 FVIII
             1 FVIIIS
           516 FVIII
                  (FVIII OR FVIIIS)
             O COMPOSITION OF FIX AND FVIII
L11
=> S FIX and treatement and haemophilia
        131131 FIX
         36375 FIXES
        155796 FIX
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(FIX OR FIXES)

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53 TREATEMENTS
          1052 TREATEMENT
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           458 HAEMOPHILIA
            13 HAEMOPHILIAS
           468 HAEMOPHILIA
                (HAEMOPHILIA OR HAEMOPHILIAS)
L12
             O FIX AND TREATEMENT AND HAEMOPHILIA
=> s FVIII and treatement and haemophilia
           516 FVIII
             1 FVIIIS
           516 FVIII
                 (FVIII OR FVIIIS)
          1002 TREATEMENT
           53 TREATEMENTS
          1052 TREATEMENT
                 (TREATEMENT OR TREATEMENTS)
           458 HAEMOPHILIA
            13 HAEMOPHILIAS
           468 HAEMOPHILIA
                 (HAEMOPHILIA OR HAEMOPHILIAS)
L13
             O FVIII AND TREATEMENT AND HAEMOPHILIA
=> s factor IX and factor VIII and haemophilia
        509873 FACTOR
        483165 FACTORS
        788251 FACTOR
                 (FACTOR OR FACTORS)
        112561 IX
             1 IXES
        112562 IX
                 (IX OR IXES)
          3425 FACTOR IX
                 (FACTOR(W)IX)
        509873 FACTOR
        483165 FACTORS
        788251 FACTOR
                 (FACTOR OR FACTORS)
        143803 VIII
            25 VIIIS
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                 (VIII OR VIIIS)
          5459 FACTOR VIII
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           458 HAEMOPHILIA
            13 HAEMOPHILIAS
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                 (HAEMOPHILIA OR HAEMOPHILIAS)
          134 FACTOR IX AND FACTOR VIII AND HAEMOPHILIA
L14
=> d rank
             1 USPATFULL
F1
=> file f1
                                                 SINCE FILE
COST IN U.S. DOLLARS
                                                                 TOTAL
                                                      ENTRY SESSION
FULL ESTIMATED COST
                                                      13.13
                                                                 42.83
FILE 'USPATFULL' ENTERED AT 11:18:21 ON 28 OCT 2004
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CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

1002 TREATEMENT

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FILE LAST UPDATED: 26 Oct 2004 (20041026/ED)
HIGHEST GRANTED PATENT NUMBER: US6810528
HIGHEST APPLICATION PUBLICATION NUMBER: US2004210975
CA INDEXING IS CURRENT THROUGH 26 Oct 2004 (20041026/UPCA)
ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 26 Oct 2004 (20041026/PD)
REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2004
USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2004
>>> USPAT2 is now available. USPATFULL contains full text of the
                                                                        <<<
>>> original, i.e., the earliest published granted patents or
                                                                        <<<
>>> applications. USPAT2 contains full text of the latest US
                                                                        <<<
>>> publications, starting in 2001, for the inventions covered in
                                                                        <<<
>>> USPATFULL. A USPATFULL record contains not only the original
>>> published document but also a list of any subsequent
>>> publications. The publication number, patent kind code, and
                                                                        <<<
>>> publication date for all the US publications for an invention
                                                                        <<<
>>> are displayed in the PI (Patent Information) field of USPATFULL
                                                                        <<<
>>> records and may be searched in standard search fields, e.g., /PN, <<<
>>> /PK, etc.
>>> USPATFULL and USPAT2 can be accessed and searched together
                                                                        <<<
>>> through the new cluster USPATALL. Type FILE USPATALL to
                                                                        <<<
>>> enter this cluster.
                                                                        <<<
>>>
                                                                        <<<
>>> Use USPATALL when searching terms such as patent assignees,
                                                                        <<<
    classifications, or claims, that may potentially change from
>>>
                                                                        <<<
>>> the earliest to the latest publication.
                                                                        <<<
This file contains CAS Registry Numbers for easy and accurate
substance identification.
=> s L14
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        483165 FACTORS
        788251 FACTOR
                 (FACTOR OR FACTORS)
        112561 IX
             1 IXES
        112562 IX
                 (IX OR IXES)
          3425 FACTOR IX
                 (FACTOR(W)IX)
        509873 FACTOR
        483165 FACTORS
        788251 FACTOR
                 (FACTOR OR FACTORS)
        143803 VIII
            25 VIIIS
        143806 VIII
                 (VIII OR VIIIS)
          5459 FACTOR VIII
                 (FACTOR(W)VIII)
           458 HAEMOPHILIA
            13 HAEMOPHILIAS
           468 HAEMOPHILIA
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L15
           134 FACTOR IX AND FACTOR VIII AND HAEMOPHILIA
=> d L14 1-1 bib, abs
L14 ANSWER 1 OF 134 USPATFULL on STN
AN
      2004:267309 USPATFULL
TI
      Mini-Ad vector for immunization
```

FILE COVERS 1971 TO PATENT PUBLICATION DATE: 26 Oct 2004 (20041026/PD)

Zhang, Wei-Wei, Libertyville, IL, UNITED STATES IN Alemany, Ramon, Grayslake, IL, UNITED STATES Dai, Yifan, Grayslake, IL, UNITED STATES Josephs, Steven, Grayslake, IL, UNITED STATES Balague, Cristina, Grayslake, IL, UNITED STATES Ayares, David, Blacksburgh, VA, UNITED STATES

Schneiderman, Richard, Highland Park, IL, UNITED STATES

US 2004208846 ÞΤ A1 20041021 A1 US 2004-837079 20040615 (9) AΙ

Continuation-in-part of Ser. No. US 1996-658961, filed on 31 May 1996, RLT ABANDONED Continuation-in-part of Ser. No. US 1997-791218, filed on 31 Jan 1997, ABANDONED

US 2000-197734P 20000418 (60) US 2000-198501P 20000418 (60) PRAI

DT Utility FS APPLICATION

MCDONNELL BOEHNEN HULBERT & BERGHOFF LLP, 300 S. WACKER DRIVE, 32ND LREP FLOOR, CHICAGO, IL, 60606

Number of Claims: 29 CLMN ECL Exemplary Claim: 1 DRWN 57 Drawing Page(s)

LN.CNT 4461

The present invention provides a method for treating a disorder such as AB hemophilia. A method of treating hemophilia in a mammal by administering recombinant virus virions comprising a nucleotide sequence having an adenoviral inverted terminal repeat fusion sequence, a packaging signal, a transcriptional control region, and a nucleic acid encoding a therapeutic protein such as FVIII. In addition, the DNA molecule does not encode an adenoviral protein. It is preferred that the virions be administered to the mammal under conditions that result in the expression of the therapeutic protein at a level that provides a therapeutic effect in said mammal.

=> d L14 1 ibib, abs

L14 ANSWER 1 OF 134 USPATFULL on STN

ACCESSION NUMBER:

2004:267309 USPATFULL

TITLE: INVENTOR(S): Mini-Ad vector for immunization

Zhang, Wei-Wei, Libertyville, IL, UNITED STATES Alemany, Ramon, Grayslake, IL, UNITED STATES Dai, Yifan, Grayslake, IL, UNITED STATES Josephs, Steven, Grayslake, IL, UNITED STATES Balague, Cristina, Grayslake, IL, UNITED STATES Ayares, David, Blacksburgh, VA, UNITED STATES

Schneiderman, Richard, Highland Park, IL, UNITED STATES

NUMBER KIND DATE -----US 2004208846 A1 20041021 US 2004-837079 A1 20040615 (9)

PATENT INFORMATION: APPLICATION INFO .: RELATED APPLN. INFO.:

Continuation-in-part of Ser. No. US 1996-658961, filed

on 31 May 1996, ABANDONED Continuation-in-part of Ser. No. US 1997-791218, filed on 31 Jan 1997, ABANDONED

NUMBER DATE -----

PRIORITY INFORMATION:

US 2000-197734P 20000418 (60) US 2000-198501P 20000418 (60)

DOCUMENT TYPE: FILE SEGMENT:

Utility APPLICATION

LEGAL REPRESENTATIVE:

MCDONNELL BOEHNEN HULBERT & BERGHOFF LLP, 300 S. WACKER DRIVE, 32ND FLOOR, CHICAGO, IL, 60606

NUMBER OF CLAIMS:

EXEMPLARY CLAIM: NUMBER OF DRAWINGS:

57 Drawing Page(s)

LINE COUNT:

4461

29

1

The present invention provides a method for treating a disorder such as hemophilia. A method of treating hemophilia in a mammal by administering recombinant virus virions comprising a nucleotide sequence having an adenoviral inverted terminal repeat fusion sequence, a packaging signal, a transcriptional control region, and a nucleic acid encoding a therapeutic protein such as FVIII. In addition, the DNA molecule does not encode an adenoviral protein. It is preferred that the virions be administered to the mammal under conditions that result in the expression of the therapeutic protein at a level that provides a therapeutic effect in said mammal.

=> s Composition and factor IX and factor VIII

760744 COMPOSITION

465872 COMPOSITIONS

818786 COMPOSITION

(COMPOSITION OR COMPOSITIONS)

509873 FACTOR

483165 FACTORS

788251 FACTOR

(FACTOR OR FACTORS)

112561 IX

1 IXES

112562 IX

(IX OR IXES)

3425 FACTOR IX

(FACTOR (W) IX)

509873 FACTOR

483165 FACTORS

788251 FACTOR

(FACTOR OR FACTORS)

143803 VIII

25 VIIIS

143806 VIII

(VIII OR VIIIS)

5459 FACTOR VIII

(FACTOR(W)VIII)

L16 1693 COMPOSITION AND FACTOR IX AND FACTOR VIII

=> d rank

F1 1 USPATFULL

=> file caplus

COST IN U.S. DOLLARS

SINCE FILE

TOTAL

FULL ESTIMATED COST

ENTRY 10.43

SESSION 53.26

FILE 'CAPLUS' ENTERED AT 11:21:35 ON 28 OCT 2004

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FILE COVERS 1907 - 28 Oct 2004 VOL 141 ISS 18 FILE LAST UPDATED: 27 Oct 2004 (20041027/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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=> s compos? and factor VIII and factor IX
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       1304812 COMPN
        522801 COMPNS
       1597236 COMPN
                  (COMPN OR COMPNS)
       2522130 COMPOS?
                  (COMPOS? OR COMPN)
        859700 FACTOR
        762249 FACTORS
       1358572 FACTOR
                 (FACTOR OR FACTORS)
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             5 VIIIS
        100451 VIII
                  (VIII OR VIIIS)
          7379 FACTOR VIII
                  (FACTOR(W)VIII)
        859700 FACTOR
        762249 FACTORS
       1358572 FACTOR
                 (FACTOR OR FACTORS)
         71711 IX
             2 IXES
         71713 IX
                  (IX OR IXES)
          3376 FACTOR IX
                  (FACTOR(W)IX)
L17
           116 COMPOS? AND FACTOR VIII AND FACTOR IX
=> d rank
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1 USPATFULL

=> file f1 COST IN U.S. DOLLARS

SINCE FILE TOTAL ENTRY SESSION FULL ESTIMATED COST 9.98 63.24

FILE 'USPATFULL' ENTERED AT 11:22:46 ON 28 OCT 2004 CA INDEXING COPYRIGHT (C) 2004 AMERICAN CHEMICAL SOCIETY (ACS)

FILE COVERS 1971 TO PATENT PUBLICATION DATE: 26 Oct 2004 (20041026/PD) FILE LAST UPDATED: 26 Oct 2004 (20041026/ED) HIGHEST GRANTED PATENT NUMBER: US6810528 HIGHEST APPLICATION PUBLICATION NUMBER: US2004210975 CA INDEXING IS CURRENT THROUGH 26 Oct 2004 (20041026/UPCA) ISSUE CLASS FIELDS (/INCL) CURRENT THROUGH: 26 Oct 2004 (20041026/PD) REVISED CLASS FIELDS (/NCL) LAST RELOADED: Aug 2004 USPTO MANUAL OF CLASSIFICATIONS THESAURUS ISSUE DATE: Aug 2004

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>>> original, i.e., the earliest published granted patents or
                                                                        <<<
>>> applications. USPAT2 contains full text of the latest US
                                                                        <<<
     publications, starting in 2001, for the inventions covered in
                                                                        <<<
    USPATFULL. A USPATFULL record contains not only the original
                                                                        <<<
     published document but also a list of any subsequent
                                                                        <<<
     publications. The publication number, patent kind code, and
                                                                        <<<
     publication date for all the US publications for an invention
                                                                        <<<
     are displayed in the PI (Patent Information) field of USPATFULL
                                                                        <<<
    records and may be searched in standard search fields, e.g., /PN, <<<
    /PK, etc.
>>>
>>> USPATFULL and USPAT2 can be accessed and searched together
                                                                        <<<
>>> through the new cluster USPATALL. Type FILE USPATALL to
                                                                        <<<
>>> enter this cluster.
                                                                        <<<
                                                                        <<<
>>> Use USPATALL when searching terms such as patent assignees,
                                                                        <<<
    classifications, or claims, that may potentially change from
                                                                        <<<
    the earliest to the latest publication.
                                                                        <<<
This file contains CAS Registry Numbers for easy and accurate
substance identification.
=> set msteps on
SET COMMAND COMPLETED
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       1344102 COMPOS?
        509873 FACTOR
        483165 FACTORS
        788251 FACTOR
                  (FACTOR OR FACTORS)
        143803 VIII
            25 VIIIS
        143806 VIII
                  (VIII OR VIIIS)
          5459 FACTOR VIII
                 (FACTOR(W)VIII)
        509873 FACTOR
        483165 FACTORS
        788251 FACTOR
                 (FACTOR OR FACTORS)
        112561 IX
             1 IXES
        112562 IX
                 (IX OR IXES)
          3425 FACTOR IX
                  (FACTOR(W)IX)
T<sub>1</sub>18
          1830 COMPOS? AND FACTOR VIII AND FACTOR IX
=> d rank
             1 USPATFULL
F1
=> d L18 1 bib,abs
L18 ANSWER 1 OF 1830 USPATFULL on STN
       2004:270005 USPATFULL
ΑN
TI
       Repressing gene expression in plants
TN
       Wu, Keqiang, Nepean, CANADA
       Miki, Brian L. A., Ottawa, CANADA
       Tian, Lining, London, CANADA
       Brown, Daniel C. W., Ilderton, CANADA
PΑ
       Her Majesty the Queen in Right of Canada, as Represented by the Minister
       of Agriculture and Agri-Food, Ottawa, CANADA (non-U.S. government)
```

PI US 6808926 B1 20041026 AI US 2000-645337 20000825 (9)

RLI Continuation-in-part of Ser. No. US 1999-383971, filed on 27 Aug 1999,

now abandoned

DT Utility FS GRANTED

EXNAM Primary Examiner: Mehta, Ashwin

LREP Oliff & Berridge, PLC CLMN Number of Claims: 38 ECL Exemplary Claim: 1

DRWN 28 Drawing Figure(s); 28 Drawing Page(s)

LN.CNT 2586

Posttranslational modification of histones, in particular acetylation and deacetylation are involved in the regulation of gene expression. Histone deacetylases remove acetyl groups from histone proteins. The present invention is directed to a method of regulating gene expression in a transgenic plant comprising, introducing into a plant a first chimeric nucleotide sequence comprising a first regulatory element in operative association with a coding sequence of interest, and an upstream activating sequence, and a second chimeric nucleotide sequence comprising a second regulatory element in operative association with a nucleotide sequence encoding histone deaceytlase and a nucleotide sequence encoding a DNA binding protein, and growing the transgenic plant. Furthermore, a method for regulating gene expression of an endogenous coding sequence of interest, or modifying a developmental, physiological or biochemical pathway in a plant is provided comprising introducing into a plant a chimeric nucleotide sequence comprising a regulatory element in operative association with a nucleotide sequence encoding histone deaceytlase fused with a nucleotide sequence encoding a DNA binding protein capable of interacting with an endogenous controlling sequence, for example an upstream activating sequence, and growing the transgenic plant. This invention also relates to novel histone deacetylase obtained from plants, to novel chimeric construct comprising these, or other histone deacetylase, and to transgenic plants, plant cells, or seeds comprising these chimeric constructs.

=> FIL CAPLUS COST IN U.S. DOLLARS

FULL ESTIMATED COST

SINCE FILE TOTAL
ENTRY SESSION
5.89 69.13

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FILE COVERS 1907 - 28 Oct 2004 VOL 141 ISS 18 FILE LAST UPDATED: 27 Oct 2004 (20041027/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

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          277963 COMPOSITIONS
          896091 COMPOSITION
                      (COMPOSITION OR COMPOSITIONS)
         1304812 COMPN
          522801 COMPNS
         1597236 COMPN
                      (COMPN OR COMPNS)
         2032719 COMPOSITION
                      (COMPOSITION OR COMPN)
          859700 FACTOR
          762249 FACTORS
         1358572 FACTOR
                      (FACTOR OR FACTORS)
          100449 VIII
                 5 VIIIS
          100451 VIII
                      (VIII OR VIIIS)
             7379 FACTOR VIII
                      (FACTOR(W)VIII)
          859700 FACTOR
          762249 FACTORS
         1358572 FACTOR
                      (FACTOR OR FACTORS)
           71711 IX
                2 IXES
           71713 IX
                      (IX OR IXES)
             3376 FACTOR IX
                      (FACTOR (W) IX)
              108 COMPOSITION AND FACTOR VIII AND FACTOR IX
L19
=> d rank
                 1
                     USPATFULL
F1
=> d L19 1 bib,abs
L19 ANSWER 1 OF 108 CAPLUS COPYRIGHT 2004 ACS on STN
      2004:817746 CAPLUS
AN
      Biologically active material conjugated with biocompatible polymer with
TI
      1:1 complex, preparation method thereof and pharmaceutical
      composition comprising the same
      Park, Myung-Ok
IN
      Biopolymed Inc., S. Korea
PA
      PCT Int. Appl., 66 pp.
SO
      CODEN: PIXXD2
DT
      Patent
LA
      English
FAN.CNT 1
                             KIND
      PATENT NO.
                                       DATE
                                                       APPLICATION NO.
                                                                                      DATE
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                                                        -----
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           2004084948 A1 20041007 WO 2004-KR701 20040327
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
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           NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE,
```

```
PRAI KR 2003-19734
                                   20030328
                            Α
     KR 2004-7983
                                   20040206
                            Α
AB
      The present invention relates to conjugates of biocompatible polymers and
     biol. active mols. wherein the activated biocompatible polymer is
      conjugated to a carboxyl group of biol. active material at a molar ratio
      of 1:1 and methods of preparation thereof and a pharmaceutical compn.
      comprising the same. Preparation of mPEG(12000)-Hz-G-CDF conjugate is
     described and its biol. activity was determined
               THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
=> s factor VIII and haemophilia
         859700 FACTOR
         762249 FACTORS
        1358572 FACTOR
                   (FACTOR OR FACTORS)
         100449 VIII
              5 VIIIS
         100451 VIII
                   (VIII OR VIIIS)
           7379 FACTOR VIII
                   (FACTOR(W)VIII)
            192 HAEMOPHILIA
              4 HAEMOPHILIAS
            195 HAEMOPHILIA
                   (HAEMOPHILIA OR HAEMOPHILIAS)
L20
            115 FACTOR VIII AND HAEMOPHILIA
=> d ranl
'RANL' IS NOT A VALID FORMAT FOR FILE 'CAPLUS'
The following are valid formats:
ABS ----- GI and AB
ALL ----- BIB, AB, IND, RE
APPS ----- AI, PRAI
BIB ----- AN, plus Bibliographic Data and PI table (default)
CAN ----- List of CA abstract numbers without answer numbers
CBIB ----- AN, plus Compressed Bibliographic Data
DALL ----- ALL, delimited (end of each field identified)
DMAX ----- MAX, delimited for post-processing
FAM ----- AN, PI and PRAI in table, plus Patent Family data FBIB ----- AN, BIB, plus Patent FAM
IND ----- Indexing data
IPC ----- International Patent Classifications
MAX ----- ALL, plus Patent FAM, RE
PATS ----- PI, SO
SAM ------ CC, SX, TI, ST, IT SCAN ----- CC, SX, TI, ST, IT (random display, no answer numbers;
               SCAN must be entered on the same line as the DISPLAY,
               e.g., D SCAN or DISPLAY SCAN)
STD ----- BIB, IPC, and NCL
IABS ----- ABS, indented with text labels
IALL ------ ALL, indented with text labels IBIB ----- BIB, indented with text labels IMAX ----- MAX, indented with text labels ISTD ----- STD, indented with text labels
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OBIB ----- AN, plus Bibliographic Data (original)

ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,

TD, TG

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OIBIB ----- OBIB, indented with text labels
SBIB ----- BIB, no citations
SIBIB ----- IBIB, no citations
HIT ----- Fields containing hit terms
HITIND ----- IC, ICA, ICI, NCL, CC and index field (ST and IT)
              containing hit terms
HITRN ----- HIT RN and its text modification
HITSTR ----- HIT RN, its text modification, its CA index name, and
              its structure diagram
HITSEQ ----- HIT RN, its text modification, its CA index name, its
              structure diagram, plus NTE and SEQ fields
FHITSTR ---- First HIT RN, its text modification, its CA index name, and
              its structure diagram
FHITSEQ ---- First HIT RN, its text modification, its CA index name, its
              structure diagram, plus NTE and SEQ fields
KWIC ----- Hit term plus 20 words on either side
OCC ----- Number of occurrence of hit term and field in which it occurs
To display a particular field or fields, enter the display field
codes. For a list of the display field codes, enter HELP DFIELDS at
an arrow prompt (=>). Examples of formats include: TI; TI, AU; BIB, ST;
TI, IND; TI, SO. You may specify the format fields in any order and the
information will be displayed in the same order as the format
specification.
All of the formats (except for SAM, SCAN, HIT, HITIND, HITRN, HITSTR,
FHITSTR, HITSEQ, FHITSEQ, KWIC, and OCC) may be used with DISPLAY ACC
to view a specified Accession Number.
ENTER DISPLAY FORMAT (BIB):
ENTER DISPLAY FORMAT (BIB):end
=> d rank
F1
             1 USPATFULL
=> s treatment of haemophilia and (factor VIII and factor IX)
       1948069 TREATMENT
       180213 TREATMENTS
      2045183 TREATMENT
                 (TREATMENT OR TREATMENTS)
           192 HAEMOPHILIA
            4 HAEMOPHILIAS
           195 HAEMOPHILIA
                 (HAEMOPHILIA OR HAEMOPHILIAS)
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                 (TREATMENT (1W) HAEMOPHILIA)
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       100451 VIII
                 (VIII OR VIIIS)
          7379 FACTOR VIII
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       762249 FACTORS
      1358572 FACTOR
                (FACTOR OR FACTORS)
         71711 IX
            2 IXES
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71713 IX

(IX OR IXES)

3376 FACTOR IX

(FACTOR(W)IX)

L21 3 TREATMENT OF HAEMOPHILIA AND (FACTOR VIII AND FACTOR IX)

=> d rank

F1 1 USPATFULL

=> d L21 1 bib, abs

L21 ANSWER 1 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2004:162533 CAPLUS

DN 140:212033

TI Non-primate lentiviral vectors for transgenic organisms preparation and gene therapy

IN Radcliffe, Philippa; Mitrophanous, Kyriacos; Themis, Michael

PA Oxford Biomedica (Uk) Limited, UK

SO U.S. Pat. Appl. Publ., 61 pp., Cont.-in-part of Appl. No. PCT/GB02/05901. CODEN: USXXCO

DT Patent

LA English

FAN.CNT 3

	PATENT NO.					KIND DATE				APPLICATION NO.					DATE			
PI	US WO WO	2004040052 2003121062 2003056022 2003056022			A1 A1 A2 A3	A1 20040226 A1 20030626 A2 20030710				US 2003-421947 US 2002-82122 WO 2002-GB5901					20020226			
	wo		AE, CO, GM, LS, PL, UA, GH, KG,	AG, CR, HR, LT, PT, UG, GM, KZ, FR,	AL, CU, HU, LU, RO, US, KE, MD, GB,	AM, CZ, ID, LV, RU, UZ, LS, RU, GR,	AT, DE, IL, MA, SC, VC, MW, TJ, IE,	2004 AU, DK, IN, MD, SD, VN, MZ, TM, IT, GN,	AZ, DM, IS, MG, SE, YU, SD, AT, LU,	DZ, JP, MK, SG, ZA, SL, BE, MC,	EC, KE, MN, SK, ZM, SZ, BG, NL,	EE, KG, MW, SL, ZW TZ, CH, PT,	ES, KP, MX, TJ, UG, CY, SE,	FI, KR, MZ, TM, ZM, CZ, SI,	GB, KZ, NO, TN, ZW, DE, SK,	GD, LC, NZ, TR, AM, DK, TR,	GE, LK, OM, TT, AZ, EE,	GH, LR, PH, TZ, BY, ES,
PRAI	GB US GB	2001-30797 2002-1140 2002-82122 2002-11409 2002-GB5901		A A A2 A		2001 2002 2002	1221 0118 0226 0517	·	ŕ	·	·	,	•					

A method of producing a transgenic cell comprising introducing into a cell AΒ a non-primate lentiviral expression vector comprising a nucleotide of interest (NOI). Also described is a method of producing a transgenic cell comprising introducing into a cell a lentiviral expression vector comprising a NOI capable of generating an antisense oligonucleotide, a ribozyme, an siRNA, a short hairpin RNA, a micro-RNA or a group 1 intron. Also described is a viral vector comprising a first nucleotide sequence, wherein said first nucleotide sequence comprises: (a) a second nucleotide sequence comprising an aptazyme; and (b) a third nucleotide sequence capable of generating a polynucleotide; wherein (a) and (b) are operably linked and wherein the aptazyme is activatable to cleave a transcript of the first nucleotide sequence such that said polynucleotide is generated. A method of producing a transgenic cell comprising introducing into a cell a non-primate lentiviral expression vector comprising a nucleotide of interest (NOI). In particular embodiments, EIAV vector expressing β -galactosidase reporter gene from an internal CMV promoter is intra-vascularly injected into perinatal mouse and detected to be expressed in various tissues. In addition, EIAV vectors expressing

corresponding therapeutic genes or ribozyme, or aptazyme (aptamer/ribozyme), or antisense RNA, or siRNA for the **treatment** of **haemophilia**, cystic fibrosis, muscular dystrophy, Parkinson's disease, and angiogenesis. Also described are production of transgenic avians as bioreactors for the production of proteins.

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=> d L21 all bib, abs
     ANSWER 1 OF 3 CAPLUS COPYRIGHT 2004 ACS on STN
     2004:162533 CAPLUS
DN
     140:212033
ED
     Entered STN: 29 Feb 2004
     Non-primate lentiviral vectors for transgenic organisms preparation and
ΤI
     gene therapy
IN
     Radcliffe, Philippa; Mitrophanous, Kyriacos; Themis, Michael
     Oxford Biomedica (Uk) Limited, UK
     U.S. Pat. Appl. Publ., 61 pp., Cont.-in-part of Appl. No. PCT/GB02/05901.
     CODEN: USXXCO
DT
     Patent
LΑ
     English
     ICM A01K067-00
IC
     ICS C12N015-867
NCL
     800021000; 435456000
     3-2 (Biochemical Genetics)
     Section cross-reference(s): 1, 12, 13, 63
FAN.CNT 3
                        KIND
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                              DATE
                                         APPLICATION NO.
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                             20040226 US 2003-421947 20030424
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    US 2004040052
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    US 2003121062
                        A1 20030626 US 2002-82122
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    WO 2003056022
                        A2
                             20030710
                                           WO 2002-GB5901
                                                                  20021223
    WO 2003056022
                        A3
                             20031231
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    WO 2003056022
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            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY,
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            FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, SI, SK, TR, BF, BJ,
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PRAI GB 2001-30797 A 20011221
                        Α
                              20020118
    GB 2002-1140
    US 2002-82122
                        A2 20020226
    GB 2002-11409
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                               20020517
    WO 2002-GB5901
                         A2
                               20021223
CLASS
PATENT NO.
             CLASS PATENT FAMILY CLASSIFICATION CODES
US 2004040052
                ICM
                       A01K067-00
                ICS
                       C12N015-867
                NCL
                       800021000; 435456000
US 2004040052
                ECLA
                       A01K067/027A; C12N015/86C
    A method of producing a transgenic cell comprising introducing into a cell
    a non-primate lentiviral expression vector comprising a nucleotide of
    interest (NOI). Also described is a method of producing a transgenic cell
    comprising introducing into a cell a lentiviral expression vector
    comprising a NOI capable of generating an antisense oligonucleotide, a
    ribozyme, an siRNA, a short hairpin RNA, a micro-RNA or a group 1 intron.
    Also described is a viral vector comprising a first nucleotide sequence,
    wherein said first nucleotide sequence comprises: (a) a second nucleotide
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sequence comprising an aptazyme; and (b) a third nucleotide sequence capable of generating a polynucleotide; wherein (a) and (b) are operably linked and wherein the aptazyme is activatable to cleave a transcript of the first nucleotide sequence such that said polynucleotide is generated. A method of producing a transgenic cell comprising introducing into a cell a non-primate lentiviral expression vector comprising a nucleotide of interest (NOI). In particular embodiments, EIAV vector expressing β -galactosidase reporter gene from an internal CMV promoter is intra-vascularly injected into perinatal mouse and detected to be expressed in various tissues. In addition, EIAV vectors expressing corresponding therapeutic genes or ribozyme, or aptazyme (aptamer/ribozyme), or antisense RNA, or siRNA for the treatment of haemophilia, cystic fibrosis, muscular dystrophy, Parkinson's disease, and angiogenesis. Also described are production of transgenic avians as bioreactors for the production of proteins. nonprimate lentivirus vector transgene delivery gene therapy Hemophilia (A; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Hemophilia (B; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Muscular dystrophy (Duchenne, gene therapy; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) CFTR (cystic fibrosis transmembrane conductance regulator) RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (EIAV vector for, for treatment of cystic fibrosis; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Dystrophin RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (EIAV vector for, for treatment of muscular dystrophy; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Transcription factors RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (HIF-1 (hypoxia-inducible factor 1), constitutive expression of, for angiogenesis treatment; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Gene, animal RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (PHD3, for angiogenesis treatment; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Gene, animal RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (PKG, promoter of; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Genetic element RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (TRE, tetracycline responsive element; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Murine leukemia virus (U3 or LTR of; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Genetic element RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (U3, of EIAV and MLV hybrid; non-primate lentiviral vectors for transgenic organisms preparation and gene therapy) Genetic element RL: BUU (Biological use, unclassified); BIOL (Biological study); USES (Uses) (U3, of EIAV; non-primate lentiviral vectors for transgenic organisms

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preparation and gene therapy)
IT
     Woodchuck hepatitis virus
         (WPRE of; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
IT
     Genetic element
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
      (Uses)
         (WPRE, wood chuck Hepatitis virus Post-transcriptional regulatory E1
        element; non-primate lentiviral vectors for transgenic organisms preparation
        and gene therapy)
     Antibodies and Immunoglobulins
     Inorganic compounds
     Nucleic acids
     Peptides, biological studies
     Proteins
     Tetracyclines
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (as aptazyme ligand, for aptazyme activation; non-primate lentiviral
        vectors for transgenic organisms preparation and gene therapy)
IT
     Embryo, animal
         (blastomere, transgenic; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IT
     Genetic element
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
      (Uses)
         (CPPT, central polypurine tract; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Digestive tract
     Egg
     Ovary
        (cell, transgenic; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IT
     Promoter (genetic element)
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (constitutive, for transgene; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
ΙT
     Peptides, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cyclic, as aptazyme ligand, for aptazyme activation; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
TТ
     Blood vessel
        (endothelium, transgene expression in; non-primate lentiviral vectors
        for transgenic organisms preparation and gene therapy)
IΤ
     Embryo, animal
        (fetus, transgenic cell in; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Hypoxia, animal
        (for VEGF-specific siRNA induction; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Post-transcriptional processing
        (gene silencing; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
     Bovine immunodeficiency virus
IT
     Caprine arthritis encephalitis virus
     Equine infectious anemia virus
     Feline immunodeficiency virus
     Human immunodeficiency virus
     Human immunodeficiency virus 1
     Visna-Maedi virus
        (gene therapy vector derived from; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Liver
        (hepatocyte, transgene expression in; non-primate lentiviral vectors
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for transgenic organisms preparation and gene therapy)
IT
     Promoter (genetic element)
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (inducible, for transgene; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Drug delivery systems
        (injections, i.m., for gene therapy vector; non-primate lentiviral
        vectors for transgenic organisms preparation and gene therapy)
ΙT
     Drug delivery systems
        (injections, i.p., for gene therapy vector; non-primate lentiviral
        vectors for transgenic organisms preparation and gene therapy)
IT
     Drug delivery systems
        (injections, i.v., for gene therapy vector; non-primate lentiviral
        vectors for transgenic organisms preparation and gene therapy)
IT
     Drug delivery systems
        (injections, intra-respiratory, for gene therapy vector; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
IT
     Drug delivery systems
        (injections, intracranial, for gene therapy vector; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
IT
     Drug delivery systems
        (injections, intrahepatic, for gene therapy vector; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
IT
     Drug delivery systems
        (injections, intraspinal, for gene therapy vector; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
IT
     Post-transcriptional processing
        (interference; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
IT
     Genetic element
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (intron, group 1; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IT
     Genetic element
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
        (long terminal repeat, EIAV hybrid; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
ΙT
     Genetic element
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (long terminal repeat, of EIAV; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Organic compounds, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (low mol. weight, as aptazyme ligand, for aptazyme activation; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
IT
    RNA
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (microRNA; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
IT
    Nerve
        (neuron, transgene expression in; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
    Adenoviral vectors
ΙT
    Angiogenesis
    Cystic fibrosis
    Gene therapy
    Molecular cloning
    Parkinson's disease
    Viral vectors
        (non-primate lentiviral vectors for transgenic organisms preparation and
        gene therapy)
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ΙT
     Antisense oligonucleotides
     Transgene
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (non-primate lentiviral vectors for transgenic organisms preparation and
        gene therapy)
IT
     Retroviral vectors
         (non-primate lentiviral; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IT
     Lentivirus
        (non-primate, gene therapy vector based on; non-primate lentiviral
        vectors for transgenic organisms preparation and gene therapy)
IT
     Egg
        (oocyte, transgenic; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IT
     Eqq
        (oogonium, cell, transgenic; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Drug delivery systems
        (oral, gastrointestinal, for gene therapy vector; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
ΙŢ
     Retroviral vectors
        (pONY8Z.1G; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
IT
     Retroviral vectors
        (pONY8Z.1ZHyb; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
ΙT
     Retroviral vectors
        (pONY8Z.4GCZ; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
IT
     Retroviral vectors
        (pONY8Z.4ZCG; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
ΙT
     Retroviral vectors
        (pONY8Z5'cppt; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
IT
     Animal cell
        (perinatal, transgenic; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IT
     Cytomegalovirus
        (promoter of; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
IT
     Aptamers
        (ribozyme hybrid, aptazyme; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     RNA
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (short hairpin RNA; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
ΙT
     Double stranded RNA
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (small interfering; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IΤ
        (spermatid, transgenic; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IΤ
        (spermatocyte, transgenic; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
        (spermatogonium, transgenic; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
        (stromal, transgenic; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
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IT
     Proteins
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (therapeutic, transgene encoding; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
     Promoter (genetic element)
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
         (tissue-specific, for transgene; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
     Ligands
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (to aptamer; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
ΙT
     Amniotic fluid
     Ascitic fluid
     Placenta
     Reproductive organ
     Umbilical cord
     Uterus
        (transgene delivery via; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
ΙT
     Ribozymes
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (transgene encoding; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
     Astrocyte
IT
     Epithelium
     Fibroblast
     Heart
     Hematopoietic precursor cell
     Kidney
     Liver
     Lung
     Lymphocyte
     Macrophage
     Monocyte
     Muscle
     Neoplasm
     Neuroglia
     Polymorphonuclear leukocyte
     Stem cell
        (transgene expression in; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IT
        (transgenic, as bioreactor; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Gamete and Germ cell
        (transgenic, gametogenesis; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     Animal
     Animal cell
     Aves
     Bos taurus
     Caenorhabditis elegans
     Drosophila
     Equus caballus
    Fish
    Human
    Insecta
    Mammalia
    Monkey
    Mus
    Oviduct
    Ovis aries
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Sperm
      Sus scrofa domestica
      Yeast
         (transgenic; non-primate lentiviral vectors for transgenic organisms
        preparation and gene therapy)
     Adeno-associated virus
     Baculoviridae
     Herpesviridae
     Parvovirus
     Poxviridae
         (vector derived from; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
IT
     Embryo, animal
         (zygote, cell, transgenic; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
IT
     109319-16-6, Factor VIII
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (EIAV vector for, for treatment of hemophilia A; non-primate lentiviral
        vectors for transgenic organisms preparation and gene therapy)
IT
     9001-28-9, Factor IX
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (EIAV vector for, for treatment of hemophilia B; non-primate lentiviral
        vectors for transgenic organisms preparation and gene therapy)
     9014-24-8, Nucleotidyltransferase, ribonucleate
     RL: BUU (Biological use, unclassified); BIOL (Biological study); USES
     (Uses)
        (II or III, promoter of gene for; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
ΙT
     50-99-7, D-Glucose, biological studies
                                              146-17-8, FMN
     Doxycycline
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (as aptazyme ligand, for aptazyme activation; non-primate lentiviral
        vectors for transgenic organisms preparation and gene therapy)
ΙT
     216864-07-2, \alpha-Synuclein
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (mutant allele, for treatment of Parkinson's disease; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
     127464-60-2, Vascular endothelial growth factor
IT
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (non-primate lentiviral vectors for transgenic organisms preparation and
        gene therapy)
ΙT
     9028-06-2, Oxygenase, protocollagen proline di-
     RL: ADV (Adverse effect, including toxicity); BSU (Biological study,
     unclassified); BIOL (Biological study)
        (ribozyme of, EIAV vector for; non-primate lentiviral vectors for
        transgenic organisms preparation and gene therapy)
     9036-22-0, Tyrosine hydroxylase 74812-49-0, Parkin
ΙT
     RL: ADV (Adverse effect, including toxicity); BSU (Biological study,
     unclassified); BIOL (Biological study)
        (ribozyme of, for treatment of Parkinson's disease; non-primate
        lentiviral vectors for transgenic organisms preparation and gene therapy)
     664350-83-8 664557-48-6 664557-49-7 664557-50-0 664557-51-1
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     664557-67-9
     RL: PRP (Properties)
        (unclaimed sequence; non-primate lentiviral vectors for transgenic
        organisms preparation and gene therapy)
     2004:162533
AN
                 CAPLUS
DN
     140:212033
     Non-primate lentiviral vectors for transgenic organisms preparation and
     gene therapy
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Reptilia

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Radcliffe, Philippa; Mitrophanous, Kyriacos; Themis, Michael
PΑ
      Oxford Biomedica (Uk) Limited, UK
      U.S. Pat. Appl. Publ., 61 pp., Cont.-in-part of Appl. No. PCT/GB02/05901.
      CODEN: USXXCO
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      Patent
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      English
FAN.CNT 3
      PATENT NO.
                           KIND DATE
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     US 2002-82122
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                                       20020226
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     WO 2002-GB5901
                                       20021223
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A method of producing a transgenic cell comprising introducing into a cell a non-primate lentiviral expression vector comprising a nucleotide of interest (NOI). Also described is a method of producing a transgenic cell comprising introducing into a cell a lentiviral expression vector comprising a NOI capable of generating an antisense oligonucleotide, a ribozyme, an siRNA, a short hairpin RNA, a micro-RNA or a group 1 intron. Also described is a viral vector comprising a first nucleotide sequence, wherein said first nucleotide sequence comprises: (a) a second nucleotide sequence comprising an aptazyme; and (b) a third nucleotide sequence capable of generating a polynucleotide; wherein (a) and (b) are operably linked and wherein the aptazyme is activatable to cleave a transcript of the first nucleotide sequence such that said polynucleotide is generated. A method of producing a transgenic cell comprising introducing into a cell a non-primate lentiviral expression vector comprising a nucleotide of interest (NOI). In particular embodiments, EIAV vector expressing β -galactosidase reporter gene from an internal CMV promoter is intra-vascularly injected into perinatal mouse and detected to be expressed in various tissues. In addition, EIAV vectors expressing corresponding therapeutic genes or ribozyme, or aptazyme (aptamer/ribozyme), or antisense RNA, or siRNA for the treatment of haemophilia, cystic fibrosis, muscular dystrophy, Parkinson's disease, and angiogenesis. Also described are production of transgenic avians as bioreactors for the production of proteins.

=> s 9001-24-8 and 109319-16-6 and composition REGISTRY INITIATED

Substance data SEARCH and crossover from CAS REGISTRY in progress... Use DISPLAY HITSTR (or FHITSTR) to directly view retrieved structures.

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0 9001-24-8
         623381 COMPOSITION
         277963 COMPOSITIONS
         896091 COMPOSITION
                  (COMPOSITION OR COMPOSITIONS)
        1304812 COMPN
        522801 COMPNS
       1597236 COMPN
                  (COMPN OR COMPNS)
       2032719 COMPOSITION
                  (COMPOSITION OR COMPN)
              0 9001-24-8 AND L23 AND COMPOSITION
L24
=> s pharmaceutical and preparation and comprising and factor VIII and factor IX
        191227 PHARMACEUTICAL
         84817 PHARMACEUTICALS
        242406 PHARMACEUTICAL
                  (PHARMACEUTICAL OR PHARMACEUTICALS)
       1312407 PREPARATION
         69780 PREPARATIONS
       1379074 PREPARATION
                  (PREPARATION OR PREPARATIONS)
       2516272 PREPN
        196827 PREPNS
       2665519 PREPN
                  (PREPN OR PREPNS)
      3393417 PREPARATION
                 (PREPARATION OR PREPN)
        326484 COMPRISING
             2 COMPRISINGS
        326485 COMPRISING
                  (COMPRISING OR COMPRISINGS)
        859700 FACTOR
        762249 FACTORS
       1358572 FACTOR
                 (FACTOR OR FACTORS)
        100449 VIII
             5 VIIIS
        100451 VIII
                 (VIII OR VIIIS)
          7379 FACTOR VIII
                 (FACTOR(W)VIII)
        859700 FACTOR
        762249 FACTORS
       1358572 FACTOR
                 (FACTOR OR FACTORS)
         71711 IX
             2 IXES
         71713 IX
                 (IX OR IXES)
          3376 FACTOR IX
                 (FACTOR (W) IX)
L25
            10 PHARMACEUTICAL AND PREPARATION AND COMPRISING AND FACTOR VIII
               AND FACTOR IX
=> d rank
             1 USPATFULL
F1
```

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L25 ANSWER 1 OF 10 CAPLUS COPYRIGHT 2004 ACS on STN
    2004:817746 CAPLUS
ED
    Entered STN: 07 Oct 2004
    Biologically active material conjugated with biocompatible polymer with
ΤI
     1:1 complex, preparation method thereof and
    pharmaceutical composition comprising the same
    Park, Myung-Ok
IN
    Biopolymed Inc., S. Korea
    PCT Int. Appl., 66 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
IC
    ICM A61K047-48
CC
    63-5 (Pharmaceuticals)
FAN.CNT 1
    PATENT NO.
                                       APPLICATION NO.
                                                            DATE
                      KIND DATE
                      A1 20041007 WO 2004-KR701
                                        -----
    WO 2004084948
                                                             20040327
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
        SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
            TD, TG
PRAI KR 2003-19734
                             20030328
                       Α
    KR 2004-7983
                             20040206
                       Α
CLASS
 PATENT NO. CLASS PATENT FAMILY CLASSIFICATION CODES
 PATENT NO.
WO 2004084948 ICM A61K047-48
    The present invention relates to conjugates of biocompatible polymers and
    biol. active mols. wherein the activated biocompatible polymer is
    conjugated to a carboxyl group of biol. active material at a molar ratio
    of 1:1 and methods of prepn. thereof and a
    pharmaceutical composition comprising the same.
    Prepn. of mPEG(12000)-Hz-G-CDF conjugate is described and its
    biol. activity was determined
    biomaterial conjugate biocompatible polymer complex prepn
st
    Agglutinins and Lectins
    Antibodies and Immunoglobulins
    Cytokines
    Enkephalins
    Growth hormone-releasing hormone receptors
    Hemoglobins
    Interleukins
    Platelet-derived growth factors
    Polymers
    Polyoxyalkylenes
    Polyphosphazenes
    Polysaccharides
    Polyurethanes
    Ricins
    Transforming growth factors
    Tumor necrosis factors
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
       (conjugates; biol. active material conjugated with biocompatible
       polymer with 1:1 complex, prepn. method thereof and
       pharmaceutical composition comprising same)
```

```
IT
      Polyamides
      RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (poly(amino acids), conjugates; biol. active material conjugated with
         biocompatible polymer with 1:1 complex, prepn. method thereof
         and pharmaceutical composition comprising same)
 IT
      Hypothalamic hormones
      RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (releasing factor, conjugates; biol. active material conjugated with
         biocompatible polymer with 1:1 complex, prepn. method thereof
         and pharmaceutical composition comprising same)
IT
      Interferons
      RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (\boldsymbol{\alpha}, \text{ conjugates}; \text{ biol. active material conjugated with}
         biocompatible polymer with 1:1 complex, prepn. method thereof
         and pharmaceutical composition comprising same)
IT
      Interferons
      RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (\beta, conjugates; biol. active material conjugated with
         biocompatible polymer with 1:1 complex, prepn. method thereof
         and pharmaceutical composition comprising same)
IT
      Interferons
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (\gamma, conjugates; biol. active material conjugated with
         biocompatible polymer with 1:1 complex, prepn. method thereof
         and pharmaceutical composition comprising same)
     9004-74-4DP, MPEG, hydrazide derivs., conjugates with biol. active mols. RL: SPN (Synthetic preparation); THU (Therapeutic use); BIOL (Biological
IT
     study); PREP (Preparation); USES (Uses)
         (biol. active material conjugated with biocompatible polymer with 1:1
         complex, prepn. method thereof and pharmaceutical
         composition comprising same)
IT
     9000-96-8D, Arginase, conjugates
                                           9001-05-2D, Catalase, conjugates
     9001-25-6D, blood coagulation factor VII, conjugates 9001-27-8D, blood
     coagulation factor VIII, conjugates 9001-28-9D,
     blood coagulation factor IX, conjugates 9001-34-7D,
     Galactosidase, conjugates 9001-37-0D, Glucose oxidase, conjugates 9001-45-0D, Glucuronidase, conjugates 9001-62-1D, Lipase, conjugates
     9002-10-2D, Tyrosinase, conjugates
                                           9002-12-4D, Uricase, conjugates
     9002-64-6D, Parathyroid hormone, conjugates
     9002-64-6D, Parathyroid hormone, conjugates 9002-71-5D, Thyroid stimulating hormone, conjugates 9002-89-5D, Polyvinyl alcohol,
                  9003-01-4D, Polyacrylic acid, conjugates
     conjugates
                                                                 9003-05-8D,
     Polyacryl amide, conjugates 9003-39-8D, Polyvinyl pyrrolidone,
                   9004-07-3D, Chymotrypsin, conjugates 9004-10-8D, Insulin,
     conjugates
                   9004-54-0D, Dextran, conjugates 9007-12-9D, Calcitonin,
     conjugates
                 9015-68-3D, Asparaginase, conjugates
     conjugates
                                                             9026-93-1D, Adenosine
     deaminase, conjugates 9027-69-4D, Adenosine diphosphatase, conjugates
     9027-98-9D, Arginine deiminase, conjugates 9033-06-1D, Glucosidase,
                  9034-40-6D, Luteinizing hormone-releasing hormone, conjugates
     conjugates
     with biocompatible polymer
                                   9054-89-1D, Superoxide dismutase, conjugates
     11096-26-7D, Erythropoietin, conjugates
                                                25104-18-1D, Poly(L-lysine),
                   25322-68-3D, Polyethylene glycol, conjugates
     conjugates
                                                                     25322-69-4D,
     Polypropyleneglycol, conjugates 26023-30-3D, Poly[oxy(1-methyl-2-oxo-1,2-
     ethanediyl)], conjugates 26100-51-6D, Polylactic acid, conjugates
     38000-06-5D, Poly(L-lysine), conjugates
                                                 62229-50-9D, Epidermal growth
     factor, conjugates 63340-72-7D, Thymic humoral factor, conjugates
     83652-28-2D, Calcitonin gene related peptide, conjugates
     Granulocyte macrophage colony stimulatingfactor, conjugates
     143011-72-7D, Granulocytecolony stimulating factor, conjugates
     345260-48-2D, Polytrimethylene glycol, conjugates
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
         (biol. active material conjugated with biocompatible polymer with 1:1
        complex, prepn. method thereof and pharmaceutical
        composition comprising same)
RE.CNT 4
              THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
```

```
(1) Enzon Inc; WO 9216555 A1 1992 CAPLUS
 (2) Enzon Inc; US 5951974 A 1999 CAPLUS
 (3) Gaertner, H; Bioconjugate Chemistry 1996, V7, P38 CAPLUS
 (4) Kirin-Amgen Inc; US 5824778 A 1998 CAPLUS
       2004:817746 CAPLUS
AN
ΤI
       Biologically active material conjugated with biocompatible polymer with
       1:1 complex, preparation method thereof and
       pharmaceutical composition comprising the same
IN
       Park, Myung-Ok
PA
       Biopolymed Inc., S. Korea
SO
       PCT Int. Appl., 66 pp.
       CODEN: PIXXD2
DT
       Patent
LA
       English
FAN.CNT 1
       PATENT NO.
                                 KIND
                                           DATE
                                                          APPLICATION NO.
                                                                                         DATE
                                ----
                                                           _____
                                            -----
           2004084948
A1 20041007 WO 2004-KR701

W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
PΤ
       WO 2004084948
                                  A1
                                           20041007 WO 2004-KR701
                                                                                          20040327
                  SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
                  TD, TG
PRAI KR 2003-19734
                                           20030328
                                   Α
       KR 2004-7983
                                  Α
                                           20040206
ΆB
       The present invention relates to conjugates of biocompatible polymers and
       biol. active mols. wherein the activated biocompatible polymer is
       conjugated to a carboxyl group of biol. active material at a molar ratio
       of 1:1 and methods of prepn. thereof and a
       pharmaceutical composition comprising the same.
       Prepn. of mPEG(12000)-Hz-G-CDF conjugate is described and its
      biol. activity was determined
=> s blood coagulation factor IX and factor VIII
         1156939 BLOOD
             1177 BLOODS
         1157059 BLOOD
                       (BLOOD OR BLOODS)
            97541 COAGULATION
              191 COAGULATIONS
            97602 COAGULATION
                       (COAGULATION OR COAGULATIONS)
          859700 FACTOR
          762249 FACTORS
         1358572 FACTOR
                      (FACTOR OR FACTORS)
            71711 IX
                 2 IXES
            71713 IX
                       (IX OR IXES)
             1840 BLOOD COAGULATION FACTOR IX
                       (BLOOD (W) COAGULATION (W) FACTOR (W) IX)
          859700 FACTOR
          762249 FACTORS
         1358572 FACTOR
                       (FACTOR OR FACTORS)
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RE

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100449 VIII
                 5 VIIIS
           100451 VIII
                       (VIII OR VIIIS)
              7379 FACTOR VIII
                       (FACTOR (W) VIII)
 L26
               480 BLOOD COAGULATION FACTOR IX AND FACTOR VIII
 => d rank
                 1
                      USPATFULL
 => d L26 1-10 bib,abs
 L26 ANSWER 1 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
       2004:817746 CAPLUS
 AN
       Biologically active material conjugated with biocompatible polymer with
 TI
       1:1 complex, preparation method thereof and pharmaceutical composition
       comprising the same
 IN
       Park, Myung-Ok
 PΑ
       Biopolymed Inc., S. Korea
       PCT Int. Appl., 66 pp.
 SO
       CODEN: PIXXD2
DT
       Patent
LA
       English
FAN.CNT 1
       PATENT NO.
                              KIND
                                                     APPLICATION NO.
                                       DATE
                                                                                  DATE
                               ----
                                        -----
                                                       -----
           004084948 A1 20041007 WO 2004-KR701 20040327
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH,
PΙ
       WO 2004084948
                CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NA, NI, NO,
                NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
           NZ, OM, PG, PH, PL, PT, RO, RO, SC, SD, SE, SG, SA, SL, SY, TO, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW

RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN,
                TD, TG
PRAI KR 2003-19734
                                Α
                                        20030328
      KR 2004-7983
                                Α
                                        20040206
AB
      The present invention relates to conjugates of biocompatible polymers and
      biol. active mols. wherein the activated biocompatible polymer is
      conjugated to a carboxyl group of biol. active material at a molar ratio
      of 1:1 and methods of preparation thereof and a pharmaceutical composition
      comprising the same. Preparation of mPEG(12000)-Hz-G-CDF conjugate is
      described and its biol. activity was determined
RE.CNT 4
                 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
                 ALL CITATIONS AVAILABLE IN THE RE FORMAT
L26
      ANSWER 2 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
AΝ
      2004:537253 CAPLUS
DN
      141:272546
TI
      Carrier detection and prenatal diagnosis on hemophilia
ΑU
      Wang, Xuefeng; Liu, Yuanfang; Liu, Xiangfan; Chu, Haiyan; Fang, Yi; Fan,
      Qishi; Wang, Hongli
CS
      Ruijin Hospital, Shanghai Second Medical University, Shanghai, 200025,
      Peop. Rep. China
      Zhonghua Jianyan Yixue Zazhi (2003), 26(9), 540-542
SO
      CODEN: ZJYZAP; ISSN: 1009-9158
PB
      Zhonghua Yixuehui Zazhishe
DT
      Journal
      Chinese
LΑ
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```
We established a simple, rapid system for carrier detection and prenatal
AB
      diagnosis on hemophilia. For hemophilia A family, factor
      VIII (FVIII) intron 22 inversion and polymorphism of
      factor VIII intragenic RFLP of Bcl I, STR within intron
      13 and 22, as well as extragenic DXS 52(ST 14) VNTR were examined by
      polymerase chain reaction; while hemophilia B family, the polymorphism of
      6 extragenic loci of factor IX (DXS1192, DXS1211, DXS8094, DXS8013,
      DXS1227, DXS102) were detected. The comprehensive utilization of direct
      assay about factor VIII intron 22 inversion and
      heredity linkage anal., the total diagnostic rate of 21 hemophilia A
      families was 94.7%; all of 10 hemophilia B families were made final
      diagnosis by using 6 extragenic loci of factor IX resp. If the intron 22
      inversion is present, we can determine the diagnosis of hemophilia A patients
      or carriers. It is a simpler and rapid method for the hemophilia carrier
      and prenatal diagnosis by detection of the intragenic and extragenic loci
     of factor VIII and extragenic loci of factor IX and
      then heredity linkage anal.
L26 ANSWER 3 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
AN
     2004:534345 CAPLUS
DN
     141:69778
TI
     Gene expression profiles in embryogenesis and marker genes for
     determination of likely success of assisted reproductive technologies
     Powers, Douglas; Wang, Shungping
IN
PA
     Embryomics, Inc., USA
SO
     PCT Int. Appl., 127 pp.
     CODEN: PIXXD2
DT
     Patent
LA
     English
FAN.CNT 1
     PATENT NO.
                        KIND
                                DATE
                                            APPLICATION NO.
                                                                     DATE
                         ----
                                              -----
PΤ
     WO 2004055217
                           A1
                                 20040701 WO 2003-US39450
                                                                      20031212
         W: AU, CA, US
         RW: AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE,
              IT, LU, MC, NL, PT, RO, SE, SI, SK, TR
PRAI US 2002-433426P
                           Ρ
                                  20021212
     Genes that show changes in patterns or levels of expression in the early
     stages of embryogenesis are identified as markers for use in assesing
     whether or not an embryo generated using assisted reproductive
     technologies is likely too implant successfully or to be carried to term.
              THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
L26 ANSWER 4 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
AN
     2004:531381 CAPLUS
DN
     141:76687
ΤI
     Stable therapeutic proteins without protease contamination produced from
     plasma or genetically engineered
IN
     Eibl, Johann
PA
     Bio-Products & Bio-Engineering Aktiengesellschaft, Austria
SO
     PCT Int. Appl., 41 pp.
     CODEN: PIXXD2
DT
     Patent
LA
    German
FAN.CNT 1
     PATENT NO.
                        KIND
                                 DATE
                                            APPLICATION NO. DATE
     WO 2004054607 A2 20040701
                                 -----
PΤ
                                             WO 2003-AT374
                                                                      20031218
        W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ,
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TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM,
               AZ, BY, KG, KZ
          RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE,
              BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU,
              MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN,
               GQ, GW, ML, MR, NE, SN, TD, TG
PRAI AT 2002-1890
                           Α
                                   20021218
      The invention concerns virus-free therapeutic proteins that are prepared
      from plasma or genetically engineered; the proteins are purified in a way
      that no enzymes, especially no proteases, zymogens are left in the product;
thus
      products with good storage stability are obtained. Proteins are isolated
      in the presence of non-toxic complexing agents, oxidation and reduction
      inhibitors, antiviral agents. Nanofiltration, cryopptn. and
      centrifugation are applied. The quality during purification is controlled by
      mass spectrometry, HPLC, gel filtration, electrophoresis and immunoassays.
      Fibrinogen-containing injections are prepared from cryoppt. with complexing
      agent and/or sodium citrate; the product contains less than 1 \mu E
      thrombin per mg fibrinogen.
     ANSWER 5 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
L26
AN
      2004:510250 CAPLUS
DN
      141:66250
     Method to produce proteins with animal glycosylation pattern in bryophyte
TI
     cells by knocking out genes for \beta 1,2-xylosyltransferase and \alpha
      1,3-fucosyltransferase and integrating human \beta 1,4-
     galactosyltransferase gene
IN
     Lienhart, Otmar
PA
     Greenovation Biotech GmbH, Germany
SO
     Eur. Pat. Appl., 47 pp.
     CODEN: EPXXDW
DT
     Patent
LΑ
     English
FAN.CNT 1
     PATENT NO.
                          KIND
                                 DATE
                                              APPLICATION NO.
                                                                       DATE
                          ----
                                  -----
                                               -----
                                 20040623 EP 2002-28536
                           A1
PΙ
     EP 1431394
                                                                       20021220
              AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
     WO 2004057002
                           A2
                                  20040708
                                               WO 2003-EP14576
     WO 2004057002
                           Α3
                                  20040826
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BW, BY, BZ, CA, CH, CN, CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, EG, ES, FI, GB, GD, GE, GH, GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR, LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO,
             NZ, OM, PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ
         RW: BW, GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE,
              BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU,
              MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN,
              GQ, GW, ML, MR, NE, SN, TD, TG
PRAI EP 2002-28536
                      Α
                               20021220
     EP 2003-22453
                           Α
                                  20031007
     Bryophyte plants, specifically Physcomitrella patens, and bryophyte plant
AΒ
     cells comprising dysfunctional fucT and xylT genes and an introduced
     glycosyl transferase gene, methods for the production of glycosylated proteins
     therewith, vectors and uses thereof. In particular, disclosed are methods
     for cloning and knocking out genes for 1,2-N-acetyl
     glucosaminyltransferase I (GNT1) , or \alpha 1,3-fucosyltransferase
     (FucT), or \beta-1,2-xylosyltransferase (XylT) from Physcomitrella
     patens; and cloning cDNA for human \beta 1,4-galactosyltransferase (GalT)
     for bryophyte cell integration. Furthermore, protoplasts derived from
     protonema of transgenic Physcomitrella plants containing human GalT and
```

FucT/XylT double knockout are transformed with human VEGF121 vector; and the detected N-glycan pattern of VEGF proteins purified in these transgenic plant is similar to that of secreted recombinant VEGF121.

RE.CNT 5 THERE ARE 5 CITED REFERENCES AVAILABLE FOR THIS RECORD
ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L26 ANSWER 6 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2004:492062 CAPLUS
- DN 141:138211
- TI The endogenous thrombin potential and high levels of coagulation factor VIII, factor IX and factor XI
- AU Siegemund, Annelie; Petros, Sirak; Siegemund, Thomas; Scholz, Ute; Seyfarth, Hans-Jurgen; Engelmann, Lothar
- CS Clinical Hemostaseology, Laboratory Practice, Medical ICU, Leipzig, Germany
- SO Blood Coagulation & Fibrinolysis (2004), 15(3), 241-244 CODEN: BLFIE7; ISSN: 0957-5235
- PB Lippincott Williams & Wilkins
- DT Journal
- LA English
- AB High plasma concns. of factor VIII, factor IX and factor XI have been reported as thrombosis risk factors. Using the thrombin generation test in platelet-poor plasma, it was aimed to describe the mechanism for this increased thrombosis risk. Endogenous thrombin potential was measured in platelet-poor plasma in 180 patients with a history of thromboembolism, and results were compared with those of 180 age-matched and sex-matched controls. Subjects with major hereditary and acquired thrombophilia were excluded. Plasma concns. of the clotting factor VIII, factor IX and factor XI were significantly elevated in patients compared with controls. The mean endogenous thrombin potential was significantly higher in patients than in controls: 191.3±3.1 (95% confidence interval, 185.3-197.4) arbitrary units vs. 180.8 \pm 2.6 (95% confidence interval, 175.7-185.9) arbitrary units (P= 0.009). The endogenous thrombin potential was significantly higher in patients with elevated factor IX and factor XI, but elevated factor VIII was not associated with a significant increase in endogenous thrombin potential. In conclusion, the increased thrombosis risk associated with high plasma concns. of factor IX and factor XI may be explained by the increase in endogenous thrombin potential. However, this did not help explain the association between elevated factor VIII and thrombosis risk.
- RE.CNT 32 THERE ARE 32 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L26 ANSWER 7 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2004:417096 CAPLUS
- DN 141:36573
- TI The role of recombinant factor VIIa (FVIIa) in fibrin structure in the absence of ${\tt FVIII/FIX}$
- AU He, S.; Blombaeck, M.; Ekman, G. Jacobsson; Hedner, U.
- CS Coagulation Research, Department of Surgical Sciences, Unit of Clinical Allergy Research, Karolinska Hospital, Karolinska Institutet, Stockholm, Swed.
- SO Journal of Thrombosis and Haemostasis (2003), 1(6), 1215-1219 CODEN: JTHOA5; ISSN: 1538-7933
- PB Blackwell Publishing Ltd.
- DT Journal
- LA English
- AB Patients with hemophilia have impaired thrombin generation and, therefore, form loose fibrin hemostatic plugs that are easily dissolved by fibrinolysis. This prevents maintained hemostasis in these patients, resulting in a severe bleeding disorder. Recombinant (F)VIIa has been shown to enhance thrombin generation on already thrombin-activated platelets in the absence of FVIII and FIX. An efficacy rate of 80-90% has

been found in hemophilia patients with inhibitors against FVIII or FIX both in association with major surgery and in the treatment of serious bleeding. In a model measuring fibrin clot permeability in a platelet-containing system described by Blomback et al. (1994), this was demonstrated to be dependent on the concentration of FVIII and FIX. The addition of

rFVIIa in concns. of 1.9, 4.8 and 9.6 μg mL-1 normalized fibrin clot permeability. The concentration of 1.9 μg mL-1 of rFVIIa normalized clot permeability in this system, and the higher concns. of rFVIIa added only slightly to the effect. No further decrease in clot permeability was found when rFVIIa in a concentration of 1.9 μg mL-1 was added to a sample with a normal concentration (100%) of FVIII or FIX. Higher concns. of rFVIIa added

the plasma containing 100% of FVIII or FIX induced only a slight further decrease of the fibrin permeability constant, arguing against any unwanted effect of extra rFVIIa on clot permeability in the case of a normal hemostasis. Furthermore, the fibrin network was studied with 3D microscopy, and the loose network found in the absence of FVIII or FIX increased in d. with increasing FVIII or FIX concns. The addition of rFVIIa to FVIII- or FIX-deficient systems altered the network structure, making the fibers thinner and more tightly packed.

RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 8 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2004:372581 CAPLUS

DN 140:387026

TI Construction of transgenic immune privileged cells for delivery of biologically active proteins and peptides and therapeutic use thereof

IN John, Constance Mary

PA USA

SO U.S. Pat. Appl. Publ., 68 pp., Cont.-in-part of U.S. Ser. No. 131,501, abandoned.

CODEN: USXXCO

DT Patent

LA English

FAN.CNT 1

	PATENT NO.	KIND	DATE	APPLICATION NO.	DATE		
PI PRAI							
	US 2004086494	A1	20040506	US 2001-941398	20010828		
	US 1996-726531	B2	19961007				
	US 1998-131501	B2	19980809				

AB The present invention provides methods for sustained delivery of biol. active proteins or peptides to mammals through immune privileged cells and therapeutic uses for nervous system diseases. Specific types of immune-privileged allogeneic or xenogenic donor cells that are naturally immune privileged are genetically modified in vitro to express or secrete the proteins or peptides. The genetically modified donor cells are subsequently implanted into host mammals and utilized for sustained delivery of biol. active proteins or peptides in vivo. The donor cells so utilized are those that inherently possess immune privilege due at least partly to the expression of Fas ligand. Methods for cell isolation, purification, tissue culture expansion, cryopreservation, gene transfer, transgene and Fas ligand expression, cell implantation, and measurement of immune responses of host animals are described.

L26 ANSWER 9 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2004:370648 CAPLUS

DN 140:380582

TI Process for sterilization of protein containing biological compositions IN Lengsfeld, Thomas; Schaefer, Wolfram; Nowak, Thomas; Grandgeorge, Michel

PA Aventis Behring GmbH, Germany

SO Eur. Pat. Appl., 15 pp.

CODEN: EPXXDW

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DT
      Patent
 LA
      English
 FAN.CNT 2
      PATENT NO.
                        KIND DATE
                                 DATE APPLICATION NO.
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 ΡI
      EP 1415669
                          A1 20040506 EP 2002-21305 20020919
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, SK
      EP 1400248
                          A1
                                20040324
                                            EP 2003-20147
                                                                    20030905
          R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
              IE, SI, LT, LV, FI, RO, MK, CY, AL, TR, BG, CZ, EE, HU, SK
     US 2004131497
                                 20040708 US 2003-663803
                         A1
                                                                     20030917
     JP 2004105740
                          A2
                                 20040408
                                             JP 2003-326061
                                                                    20030918
PRAI EP 2002-21305
                          Α
                                 20020919
     The present invention relates to a method for inactivating microorganisms
     especially viruses in protein containing biol. compns., especially blood,
placental, serum
     and plasma components or derivs. solns. of human or animal origin or
     compns. containing proteins obtained by the extraction of vegetal or animal
tissues
     or obtained by biotechnol. techniques, i.e. by culture of natural or
     recombinant cells of bacterial, yeast, plant or human or animal origin
     thereby retaining the integrity of the desired protein at a degree
     suitable for its purpose. By extension the method applies to the
     inactivation of bacterial or viral prepns., when protein components or
     protein antigens are present, which are intended for use in inactivated or
     non-living vaccines. E.g., fibrinogen was stabilized with rutin or
     vanillin before sterilization by UV irradiation
RE.CNT 11
              THERE ARE 11 CITED REFERENCES AVAILABLE FOR THIS RECORD
              ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 10 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
AN
     2004:331821 CAPLUS
DN
     140:353209
     Glycan remodeling and glycoconjugation of granulocyte colony stimulating
TI
     factor
     Defrees, Shawn; Zopf, David; Bayer, Robert; Bowe, Caryn; Hakes, David;
IN
     Chen, Xi
     Neose Technologies, Inc., USA
PΑ
     U.S. Pat. Appl. Publ., 754 pp., Cont.-in-part of U.S. Ser. No. 360,779.
SO
     CODEN: USXXCO
DT
     Patent
LA
     English
FAN.CNT 12
                    KIND DATE APPLICATION NO. DATE
     PATENT NO.
     US 2004077836 A1 20040422 US 2003-410962
WO 2003031464 A2 20030417 WO 2002-US32263
ΡI
                                                                    20030409
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
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             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
             UA, UG, US, UZ, VN, YU, ZA, ZM, ZW
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             SZ, SZ, TZ, TZ, UG, UG, ZM, ZM, ZW, ZW, AM, AZ, BY, KG, KZ, MD,
             RU, TJ, TM, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB,
             GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM,
             GA, GN, GQ
    US 2004137557 A1 20040715

US 2001-328523P P 20011010

US 2001-344692P P 20011019

US 2001-334233P P 20011128

US 2001-334301P P 20011128
                                20040715
                                            US 2002-287994
                                                                    20021105
PRAI US 2001-328523P
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US 2002-387292P
                     Р
                           20020607
US 2002-391777P
                          20020625
                     P
                         20020717
20020816
20020828
US 2002-396594P
                     Ρ
US 2002-404249P
                     P
US 2002-407527P
                     Р
WO 2002-US32263
                     A1
                          20021009
US 2002-287994
                     A2
                           20021105
US 2003-360770
                     A2
                           20030106
US 2003-360779
                     A2
                           20030219
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The invention includes a multitude of methods of remodeling a peptide to AB have a specific glycan structure attached to the peptide. The methods comprise cell-free in vitro addition and/or deletion of sugars to or from a peptide mol. in such a manner as to provide a glycopeptide mol. having a specific customized or desired glycosylation pattern, wherein the glycopeptide is produced at an industrial scale and is suitable for therapeutic use in a mammal. The modified sugar that has been added to the peptide is generated via an enzymic reaction, because enzyme-based addition of conjugate mols. to peptides has the advantage of regioselectivity and stereoselectivity. Thus, a granulocyte colony stimulating factor (G-CSF) peptide that is expressed in a mammalian cell system is trimmed back using a sialidase. The residues thus exposed are modified by the addition of a sialic acid-poly(ethylene glycol) moiety, using an appropriate donor therefor and ST3Gall. Mammalian cell expressed G-CSF is contacted with a sialic acid donor that is modified with levulinic acid, adding a reactive ketone to the sialic acid donor. After addition to a glycosyl residue on the glycan on the peptide, the ketone is derivatized with a moiety such as hydrazine- or amino-PEG. Analogous schemes are provided for G-CSF expressed in an insect or bacterial cell.

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L4

L5

(FILE 'HOME' ENTERED AT 11:02:09 ON 28 OCT 2004)

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 11:02:32 ON 28 OCT 2004 SEA COMPOSIT? AND FACTOR IX AND FACROR VIII

> 1 FILE USPATFULL QUE COMPOSIT? AND FACTOR IX AND FACROR VIII

FILE 'USPATFULL' ENTERED AT 11:05:46 ON 28 OCT 2004

 L_2 1 S L1

L3 0 S L1 AND HAEMOPHILIA

171 S COMPOSITION AND FIX AND FVIII

0 S FVIII AND TREATEMENT AND HAEMOPHILIA

FILE 'CAPLUS' ENTERED AT 11:11:41 ON 28 OCT 2004 L6 3 S FACTOR FVIII AND FIX

FILE '1MOBILITY' ENTERED AT 11:12:41 ON 28 OCT 2004

FILE 'USPATFULL' ENTERED AT 11:12:54 ON 28 OCT 2004 1 S L1

L7 L_8 1 S L7

FILE 'USPATFULL' ENTERED AT 11:13:59 ON 28 OCT 2004

L9 1 S L8 AND COMPOSITION L10

0 S L9 AND HAEMOPHILIA

L110 S COMPOSITION OF FIX AND FVIII

L120 S FIX AND TREATEMENT AND HAEMOPHILIA

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L13
                   0 S FVIII AND TREATEMENT AND HAEMOPHILIA
 L14
                134 S FACTOR IX AND FACTOR VIII AND HAEMOPHILIA
       FILE 'USPATFULL' ENTERED AT 11:18:21 ON 28 OCT 2004
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                134 S L14
 L16
               1693 S COMPOSITION AND FACTOR IX AND FACTOR VIII
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 L17
                116 S COMPOS? AND FACTOR VIII AND FACTOR IX
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 L18
              1830 S L17
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 T<sub>1</sub>19
               108 S COMPOSITION AND FACTOR VIII AND FACTOR IX
                115 S FACTOR VIII AND HAEMOPHILIA
 L20
                  3 S TREATMENT OF HAEMOPHILIA AND (FACTOR VIII AND FACTOR IX)
 L21
                    S 9001-24-8 AND 109319-16-6/REG# AND COMPOSITION
       FILE 'REGISTRY' ENTERED AT 11:30:13 ON 28 OCT 2004
L22
               1 S 109319-16-6/RN
       FILE 'CAPLUS' ENTERED AT 11:30:14 ON 28 OCT 2004
L23
              4039 S L22
L24
                 0 S 9001-24-8 AND L23 AND COMPOSITION
L25
                10 S PHARMACEUTICAL AND PREPARATION AND COMPRISING AND FACTOR VIII
L26
               480 S BLOOD COAGULATION FACTOR IX AND FACTOR VIII
=> s L26 and haemophilia
              192 HAEMOPHILIA
                4 HAEMOPHILIAS
              195 HAEMOPHILIA
                    (HAEMOPHILIA OR HAEMOPHILIAS)
L27
                7 L26 AND HAEMOPHILIA
=> d L26 10-20 bib,abs
L26 ANSWER 10 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
AN
      2004:331821 CAPLUS
DN
      140:353209
      Glycan remodeling and glycoconjugation of granulocyte colony stimulating
ΤI
      factor
      Defrees, Shawn; Zopf, David; Bayer, Robert; Bowe, Caryn; Hakes, David;
IN
      Chen, Xi
PA
      Neose Technologies, Inc., USA
      U.S. Pat. Appl. Publ., 754 pp., Cont.-in-part of U.S. Ser. No. 360,779.
SO
      CODEN: USXXCO
DT
      Patent
LΑ
     English
FAN.CNT 12
                            KIND DATE APPLICATION NO. DATE
      PATENT NO.
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      US 2004077836
      A1 20040422
      US 2003-410962
      20030409

      WO 2003031464
      A2 20030417
      WO 2002-US32263
      20021009

PΤ
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GR, IE, IT, LU, MC, NL, PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM,
              GA, GN, GQ
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                                                 US 2002-287994
                                                                           20021105
PRAI US 2001-328523P
                             Ρ
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     US 2001-344692P
                            P
                                   20011019
     US 2001-334233P
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                                  20011128
                        . Р
     US 2001-334301P
                                  20011128
     US 2002-387292P
                            P
                                  20020607
     US 2002-391777P
                            P
     US 2002-396594P

US 2002-404249P

US 2002-407527P

WO 2002-US32263

US 2002-287994

US 2003-360770
                                  20020625
                            P
                                  20020717
                           P
                                 20020816
                            P
                                  20020828
                            A1
                                   20021009
                            A2
                                   20021105
                            A2
                                   20030106
     US 2003-360779
                            A2
                                   20030219
     The invention includes a multitude of methods of remodeling a peptide to
AB
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have a specific glycan structure attached to the peptide. The methods comprise cell-free in vitro addition and/or deletion of sugars to or from a peptide mol. in such a manner as to provide a glycopeptide mol. having a specific customized or desired glycosylation pattern, wherein the glycopeptide is produced at an industrial scale and is suitable for therapeutic use in a mammal. The modified sugar that has been added to the peptide is generated via an enzymic reaction, because enzyme-based addition of conjugate mols. to peptides has the advantage of regionelectivity and stereoselectivity. Thus, a granulocyte colony stimulating factor (G-CSF) peptide that is expressed in a mammalian cell system is trimmed back using a sialidase. The residues thus exposed are modified by the addition of a sialic acid-poly(ethylene glycol) moiety, using an appropriate donor therefor and ST3Gall. Mammalian cell expressed G-CSF is contacted with a sialic acid donor that is modified with levulinic acid, adding a reactive ketone to the sialic acid donor. After addition to a glycosyl residue on the glycan on the peptide, the ketone is derivatized with a moiety such as hydrazine- or amino-PEG. Analogous schemes are provided for G-CSF expressed in an insect or bacterial cell.

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ANSWER 11 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
L26
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AΝ 2004:306636 CAPLUS

DN141:293779

Different thrombotic risk factors - contribution to the endogenous TIthrombin potential ΑU

Siegemund, A.; Siegemund, T.; Scholz, U.; Petros, S.; Engelmann, L.

CS Germany

Hemophilia Symposium, 33rd, Hamburg, Germany, 2002 (2004), Meeting Date SO 2002, 257-261. Editor(s): Scharrer, Inge; Schramm, Wolfgang. Publisher: Springer-Verlag, Berlin, Germany. CODEN: 69FGWY; ISBN: 3-540-00902-7

DT Conference

LΑ English

In 563 patients with thrombosis, several risk factors were measured, AΒ including PC-resistance (in cases of lowered ratio Factor V Leiden), prothrombin level and prothrombin mutation 20210 GA, the coagulation factors VIII:C, IX and XI, protein C, protein S, and antithrombin. A strong correlation was found between the number of thrombotic risk factors and the amount of generated thrombin. One risk factor alone was associated with a normal endogenous thrombin potential (ETP), but two risk factors or more resulted in an increase of ETP. Not all thrombogenic risk factors contributed in the same manner to the ETP. There was a simultaneous increase in ETP with higher levels of coagulation factors indicating a continuous dose-response relation between ETP. Elevated levels of FVIII:C and Factor V-Leiden did not influence the ETP. In contrast, the simultaneous occurrence of Factor V-Leiden and prothrombin allele 20210 GA decreased the ETP.

THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD RE.CNT 10

ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L26 ANSWER 12 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
- AN2004:306620 CAPLUS
- Endogenous thrombin potential in platelet-rich plasma. New insights TΤ regarding the different action of FVIII and FIX
- Siegemund, A.; Siegemund, T.; Scholz, U.; Petros, S.; Engelmann, L. ΑU
- CS Germany
- Hemophilia Symposium, 33rd, Hamburg, Germany, 2002 (2004), Meeting Date SO 2002, 87-93. Editor(s): Scharrer, Inge; Schramm, Wolfgang. Publisher: Springer-Verlag, Berlin, Germany. CODEN: 69FGWY; ISBN: 3-540-00902-7
- DTConference
- LAEnglish
- The role of platelets in the thrombin generation process was investigated. AB Endogenous thrombin potential (ETP) was measured with some modifications in substrate and activator concns. Measurement of thrombin generation is gaining an increasing importance in the field of hemostaseol. The ETP demonstrates the overall potential of the hemostatic system and represents the interactions between coagulation factors and platelets. The results confirm that the major defect in hemophilia A and B is the decreased ability to generate enough thrombin. The reduced thrombin generation is more obvious in hemophilia B than A. Data showed a high FVIII-related thrombin generation in hemophilia A and a high platelet-related thrombin generation in hemophilia B. Data emphasized the cell based model of thrombin generation. Measuring thrombin generation in platelet-rich plasma is a good method to describe such interaction between coagulation factors and platelets.
- RE.CNT 4 THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- ANSWER 13 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
- AN2004:269906 CAPLUS

US 2002-387292P

P

- DN 140:300039
- Remodeling of protein-linked oligosaccharide moieties and the resulting TIglycoproteins and glycopeptides
- Defree, Shawn; Zopf, David; Bayer, Robert; Hakes, David; Chen, Xi IN
- PΑ Neose Technologies Inc., USA
- U.S. Pat. Appl. Publ., 749 pp., Cont.-in-part of U.S. Ser. No. 360,779. SO CODEN: USXXCO
- DTPatent
- LA English
- FAN.CNT 12

FAN.	CNT	12																	
	PA 	ATENT NO.				KIN	D -	DATE			APPL	ICAT	ION	NO.		D	ATE		
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PRAI	US US US	2004 2001 2001 2001 2001	13755 -3285 -3446 -3342	57 523P 592P 233P	GQ	A1 P P P		20040 2001: 2001: 2001: 2001:	0715 1010 1019 1128										

20020607

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US 2002-391777P
                              P
                                       20020625
       US 2002-396594P
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                                       20020717
       US 2002-404249P
                              P
                                       20020816
       US 2002-407527P
                              P
                                      20020828
       WO 2002-US32263
                              A1
                                      20021009
       US 2002-287994
                               A2
                                      20021105
       US 2003-360770
                               A2
                                      20030106
      US 2003-360779
                              A2
                                      20030219
      The invention includes a multitude of methods of remodeling a peptide to
      have a specific glycan structure attached to the peptide. The methods
      comprise cell-free in vitro addition and/or deletion of sugars to or from a
      peptide mol. in such a manner as to provide a glycopeptide mol. having a
      specific customized or desired glycosylation pattern, wherein the
      glycopeptide is produced at an industrial scale. The modified sugar that
      has been added to the peptide is generated via an enzymic reaction,
      because enzyme-based addition of conjugate mols. to peptides has the
      advantage of regioselectivity and stereoselectivity. A key feature of the
      invention is to take a peptide produced by any cell type and generate a
      core glycan structure on the peptide, following which the glycan structure
      is then remodeled in vitro to generate a glycopeptide having a
      glycosylation pattern suitable for therapeutic use in a mammal.
L26 ANSWER 14 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
      2004:204052 CAPLUS
      140:213540
      Diagnostic assay for thrombin-activatable fibrinolysis inhibitor (TAFI)
      Greenfield, Robert S.; An, Seong Soo A.
      American Diagnostica, Inc., USA
      PCT Int. Appl., 37 pp.
      CODEN: PIXXD2
      Patent
      English
FAN.CNT 1
      PATENT NO.
                           KIND DATE
                                                 APPLICATION NO.
                                                                             DATE
                            ____
                                     -----
                                                   -----
          2004020976 A2 20040311 WO 2003-US27061 20030829
2004020976 A3 20040812
W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NI, NO, NZ, OM,
BC, BH, BI, DT, BO, RH, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN,
      WO 2004020976
      WO 2004020976
               PG, PH, PL, PT, RO, RU, SC, SD, SE, SG, SK, SL, SY, TJ, TM, TN,
               TR, TT, TZ, UA, UG, US, UZ, VC, VN, YU, ZA, ZM, ZW, AM, AZ, BY,
               KG, KZ, MD, RU
          RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRAI US 2002-406756P
                            P
                                 20020829
     The invention relates to a diagnostic assay for selectively measuring
     levels of the 35kD form of thrombin-activatable fibrinolysis inhibitor
     (TAFIa or TAFIai), or a derivative or variant thereof, but not the TAFI
     proenzyme (TAFI) or the N-terminal activation peptide of TAFI.
    ANSWER 15 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
     2004:62593 CAPLUS
     141:813
     Long-term administration of highly purified eicosapentaenoic acid ethyl
     ester improves blood coagulation abnormalities and dysfunction of vascular
     endothelial cells in Otsuka Long-Evans Tokushima Fatty rats
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AΒ

DN

TIIN

PASO

DT

LA

PΙ

L26

ANDN

ΤI

ΑU

Tajima, Naoko Department of Internal Medicine, National Higashi-Utsunomiya Hospital, CS Kawachi-machi, Kawachi-gun, Tochigi, 329-1193, Japan

Mori, Yutaka; Nobutaka, Hidefumi; Harada, Tsuyoshi; Kasahara, Toshihiko;

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Endocrine Journal (Kyoto, Japan) (2003), 50(5), 603-611
 SO
      CODEN: ENJOEO; ISSN: 0918-8959
 PΒ
      Japan Endocrine Society
 DT
      Journal
 LA
      English
      We investigated the effect of highly purified eicosapentaenoic acid Et
 AB
      ester (EPA-E) on blood coagulation abnormalities and dysfunction of
      vascular endothelial cells in spontaneously diabetic Otsuka Long-Evans
      Tokushima Fatty rats. The animals were treated with either EPA-E or lard
      at a daily dose of 0.3 g/kg/day for 52 wk by gavage, and their
      coagulation/fibrinolytic parameters, platelet aggregation, and functions
      of the vascular endothelial cells were examined EPA-E significantly
      improved coagulation-related parameters including prothrombin time,
      activated partial thromboplastin time, fibrinogen level, and activities of
      factor II, V, VII, VIII, IX, X, XI, and XII, and antithrombin III, and
      fibrinolysis-related parameters including plasminogen, tissue-type
      plasminogen activator, \alpha 2-plasmin inhibitor, and plasminogen
      activator inhibitor. It also suppressed ADP- or collagen-induced platelet
      aggregation and the cholesterol/phospholipid molar ratio in platelet
      membranes at a dose of 0.3 g/kg. In addition, it significantly increased the
      migration activity of vascular endothelial cells, and decreased the
      binding of vascular endothelial cells to vascular endothelial growth
      factor. In contrast, lard had no effect on hypercoagulation,
     hypofibrinolysis, and platelet hyperaggregation but significantly
      aggravated the dysfunction of vascular endothelial cells. These data
     demonstrate that EPA-E beneficially altered certain factors known to
     promote thrombosis and atherosclerosis in this animal model.
RE.CNT 10
               THERE ARE 10 CITED REFERENCES AVAILABLE FOR THIS RECORD
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
     ANSWER 16 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
     2004:19771 CAPLUS
DN
     140:105280
     Thrombin-cleavable factor X analogs, and use for procoagulants
TI
     Louvain, Virginie; Bianchini, Elsa; Marque, Pierre Emmanuel; Calmel,
     Tareau Claire; Aiach, Martine; Le Bonniec, Bernard
     Institut National de la Sante et de la Recherche Medicale INSERM, Fr.
PA
     Fr. Demande, 63 pp.
SO
     CODEN: FRXXBL
DT
     Patent
LA
     French
FAN.CNT 1
     PATENT NO.
                        KIND
                                DATE
                                           APPLICATION NO.
                                                                   DATE
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PΙ
     FR 2841904
                          A1
                                 20040109
                                            FR 2002-8299
                                                                     20020703
     FR 2841904
                          В1
                                 20040820
         WO 2004005347
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             KG, KZ, MD, RU
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG, CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, HU, IE, IT, LU, MC, NL, PT, RO, SE, SI, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
PRAI FR 2002-8299
                          A
                                20020703
    MARPAT 140:105280
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AB The invention discloses analogs of factor X having a thrombin-cleavable sequence Pro-Arg-Ala in place of the sequence Thr-Arg-Ile at the activation site of native factor X. These factor X analogs are useful for obtaining procoagulant drugs.

RE.CNT 1 THERE ARE 1 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L26 ANSWER 17 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2003:1013268 CAPLUS
- DN 140:230875
- TI The effects of different alcoholic drinks on lipids, insulin and haemostatic and inflammatory markers in older men
- AU Wannamethee, Sasiwarang Goya; Lowe, Gordon D. O.; Shaper, Gerald; Whincup, Peter H.; Rumley, Ann; Walker, Mary; Lennon, Lucy
- CS Department of Primary Care and Population Sciences, Royal Free and University College Medical School, London, UK
- SO Thrombosis and Haemostasis (2003), 90(6), 1080-1087 CODEN: THHADQ; ISSN: 0340-6245
- PB Schattauer GmbH
- DT Journal
- LA English
- Light to moderate drinking is associated with lower risk of coronary heart (CHD) than non-drinkers. We have examined the relationships between total alc. intake and type of alc. beverage and several potential biol. mechanisms. We carried out the study in 3158 men aged 60-79 yr drawn from general practices in 24 British towns with no history of myocardial infarction, stroke or diabetes and who were not on warfarin. Total alc. consumption showed a significant pos. dose-response relationship with high d. lipoprotein cholesterol (HDL-C), coagulation factor IX, haematocrit, blood viscosity, and tissue plasminogen antigen, and an inverse dose-response relationship with insulin, fibrinogen, von Willebrand factor (vWF) and triglycerides after adjustment for possible confounders. Total alc. consumption showed weak assocns. with plasma viscosity and fibrin D-dimer, and no association with factors VII, VIII, or C-reactive protein (CRP). Wine was specifically associated with lower CRP, plasma viscosity, factor VIII and triglycerides. The findings are consistent with the suggestion that HDL-C in particular but also insulin and haemostatic factors may contribute to the beneficial effect of light to moderate drinking on risk of CHD. Wine has effects that may confer greater protection than other alc. beverages.
- RE.CNT 36 THERE ARE 36 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L26 ANSWER 18 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2003:1000795 CAPLUS
- DN 141:4643
- TI Discontinuous residues of factor IX constitute a surface for binding the anti-factor IX monoclonal antibody A-5
- AU Chang, Yu-Jia; Wu, Hua-Lin; Hsu, Ya-Chu; Hamaguchi, Nobuko; Shi, Guey-Yueh; Shen, Ming-Ching; Lin, Shu-Wha
- CS College of Medicine, Institute of Basic Medical Sciences, National Cheng Kung University, Tainan, Taiwan
- SO Thrombosis Research (2003), 111(4-5), 293-299 CODEN: THBRAA; ISSN: 0049-3848
- PB Elsevier Science Inc.
- DT Journal
- LA English
- Anti-human factor IX monoclonal antibody, A-5 (Mab A-5), has been widely used in structure-function studies for factor IX. Mab A-5 recognizes the catalytic domain of human factor IX (FIX). Regions important for Mab A-5 binding have previously been localized to the amino terminus of the heavy chain of factor IX, encompassing amino acid residues 181-310 [Blood (74) 971]. We have selected 20 positions in this region for alanine-scanning mutagenesis. We found that Mab A-5 failed to react with the recombinant factor IX mutants with substitutions at positions 228 and 252. Mab A-5 also reacted to a lesser extent to FIXD276A (factor IX with alanine substitution for aspartic acid at residue 276) and FIXK201A/D203A (double alanine substitutions at residues 201 and 203). The apparent dissociation

rate consts. (KD) in binding Mab A-5 were 6.0+10-9, 1.4+10-8 and 2.0+10-8 M, for wild-type FIX, FIXK201A/D203A and FIXD276A, resp. The increased KD values of the two FIX mutants are mainly owing to the increased dissociation rates. These affected residues constitute a surface opposite from the factor VIIIa binding surface. We conclude that the epitope for Mab A-5 is on a surface composed of residues 228, 252, 276, and 201 or 203. This surface, which may not be important for factor VIII binding, contains a strong antigenic region on factor IX.

RE.CNT 27 THERE ARE 27 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L26 ANSWER 19 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2003:987097 CAPLUS

- DN 140:263538
- TI Clotting factors VIII and IX
- AU Brownlee, George G.; Giangrande, Paul L. F.
- CS Chemical Pathology Unit, Sir William Dunn School of Pathology, University of Oxford, Oxford, UK
- SO Recombinant Protein Drugs (2001), 67-88. Editor(s): Buckel, Peter. Publisher: Birkhaeuser Verlag, Basel, Switz. CODEN: 69EWGR; ISBN: 3-7643-5904-8
- DT Conference; General Review
- LA English
- AB A review on recombinant **factors VIII** and IX for the treatment of patients with hemophilia A and B.
- RE.CNT 37 THERE ARE 37 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT
- L26 ANSWER 20 OF 480 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2003:975512 CAPLUS
- DN 140:35203
- TI Immunogenicity and immune tolerance coagulation **factors VIII** and IX
- AU Rup, B.
- CS Bioanalytical Research & Development, Wyeth Research, Andover, MA, USA
- Developments in Biologicals (Basel, Switzerland) (2003), 112 (Immunogenicity of Therapeutic Biological Products), 55-59 CODEN: DBEIAI; ISSN: 1424-6074
- PB S. Karger AG
- DT Journal; General Review
- LA English
- AF A review. Some of the major issues related to the development and control of antibodies that occur during treatment of hemophilia with replacement factors (Factor VIII and Factor IX) are reviewed.

 Information on anal. issues, immunogenicity, and immune tolerance may be applicable to the study of other therapeutic proteins. Conversely, new information obtained from evaluation of other therapeutic protein products may address issues that remain unresolved for Factor VIII and FIX replacement therapy.
- RE.CNT 19 THERE ARE 19 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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(FILE 'HOME' ENTERED AT 11:02:09 ON 28 OCT 2004)

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 11:02:32 ON 28 OCT 2004 SEA COMPOSIT? AND FACTOR IX AND FACROR VIII

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L1
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 L2
               1 S L1
 L3
               0 S L1 AND HAEMOPHILIA
 L4
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 L5
               0 S FVIII AND TREATEMENT AND HAEMOPHILIA
      FILE 'CAPLUS' ENTERED AT 11:11:41 ON 28 OCT 2004
 L6
               3 S FACTOR FVIII AND FIX
      FILE '1MOBILITY' ENTERED AT 11:12:41 ON 28 OCT 2004
      FILE 'USPATFULL' ENTERED AT 11:12:54 ON 28 OCT 2004
 L7
               1 S L1
 L8
               1 S L7
     FILE 'USPATFULL' ENTERED AT 11:13:59 ON 28 OCT 2004
 L9
               1 S L8 AND COMPOSITION
 L10
               0 S L9 AND HAEMOPHILIA
 L11
               0 S COMPOSITION OF FIX AND FVIII
 L12
               0 S FIX AND TREATEMENT AND HAEMOPHILIA
L13
              0 S FVIII AND TREATEMENT AND HAEMOPHILIA
L14
            134 S FACTOR IX AND FACTOR VIII AND HAEMOPHILIA
     FILE 'USPATFULL' ENTERED AT 11:18:21 ON 28 OCT 2004
L15
            134 S L14
L16
           1693 S COMPOSITION AND FACTOR IX AND FACTOR VIII
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L17
            116 S COMPOS? AND FACTOR VIII AND FACTOR IX
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                SET MSTEPS ON
L18
           1830 S L17
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L19
            108 S COMPOSITION AND FACTOR VIII AND FACTOR IX
L20
            115 S FACTOR VIII AND HAEMOPHILIA
L21
              3 S TREATMENT OF HAEMOPHILIA AND (FACTOR VIII AND FACTOR IX)
                S 9001-24-8 AND 109319-16-6/REG# AND COMPOSITION
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L22
              1 S 109319-16-6/RN
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L23
           4039 S L22
L24
              0 S 9001-24-8 AND L23 AND COMPOSITION
            10 S PHARMACEUTICAL AND PREPARATION AND COMPRISING AND FACTOR VIII
L25
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L26
L27
             7 S L26 AND HAEMOPHILIA
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          191 COAGULATIONS
         97602 COAGULATION
                 (COAGULATION OR COAGULATIONS)
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FILE USPATFULL

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1 S L1
 L3
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 L_4
             171 S COMPOSITION AND FIX AND FVIII
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      FILE 'CAPLUS' ENTERED AT 11:11:41 ON 28 OCT 2004
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      FILE '1MOBILITY' ENTERED AT 11:12:41 ON 28 OCT 2004
      FILE 'USPATFULL' ENTERED AT 11:12:54 ON 28 OCT 2004
 L7
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     FILE 'USPATFULL' ENTERED AT 11:13:59 ON 28 OCT 2004
L9
               1 S L8 AND COMPOSITION
L10
               0 S L9 AND HAEMOPHILIA
L11
               0 S COMPOSITION OF FIX AND FVIII
L12
              0 S FIX AND TREATEMENT AND HAEMOPHILIA
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L14
            134 S FACTOR IX AND FACTOR VIII AND HAEMOPHILIA
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           1693 S COMPOSITION AND FACTOR IX AND FACTOR VIII
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L19
            108 S COMPOSITION AND FACTOR VIII AND FACTOR IX
L20
            115 S FACTOR VIII AND HAEMOPHILIA
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             10 S PHARMACEUTICAL AND PREPARATION AND COMPRISING AND FACTOR VIII
L26
            480 S BLOOD COAGULATION FACTOR IX AND FACTOR VIII
             7 S L26 AND HAEMOPHILIA
L27
            425 S BLOOD COAGULATION FACTOR VIII AND FACTOR IX AND PY<2003
L28
L29
              6 S L28 AND HAEMOPHILIA
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         1177 BLOODS
       1157059 BLOOD
                 (BLOOD OR BLOODS)
         97541 COAGULATION
           191 COAGULATIONS
         97602 COAGULATION
                 (COAGULATION OR COAGULATIONS)
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       762249 FACTORS
       1358572 FACTOR
                 (FACTOR OR FACTORS)
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 100451 VIII
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  11554 FIX
   2307 FIXES
  13733 FIX
          (FIX OR FIXES)
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 277963 COMPOSITIONS
 896091 COMPOSITION
          (COMPOSITION OR COMPOSITIONS)
1304812 COMPN
522801 COMPNS
1597236 COMPN
          (COMPN OR COMPNS)
2032719 COMPOSITION
          (COMPOSITION OR COMPN)
    192 HAEMOPHILIA
      4 HAEMOPHILIAS
    195 HAEMOPHILIA
          (HAEMOPHILIA OR HAEMOPHILIAS)
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0 BLOOD COAGULATION FACTOR VIII AND FIX AND COMPOSITION AND HAEMOP

L30

HILIA

Factor VIII (FVIII) or Factor IX

(FIX) can be well controlled with periodic iv. injections of FVIII or FIX concs. Either concentrate can be isolated from large human pools (i.e., plasma-derived FVIII or FIX concentrate) or from culture supernatants of recombinant cells engineered to secrete FVIII or FIX. The validated viral inactivation strategies used by manufacturers of FVIII and FIX concs. have essentially eliminated the transmission of hepatitis B, hepatitis C and HIV viruses. The low yields and inherent instability of FVIII (and FVIIIa in particular) and the addnl. costs of viral inactivation methods make the annual cost/patient for prophylaxis and treatment of hemophilia very expensive. Several strategies have been adopted and proposed to improve yields of FVIII. These include: deletion of portions of FVIII which are not associated with function; mutations to prevent inactivation of FVIII by protease degradation; and synthesis of FVIII fragments to replace portions deleted in some FVIII deficient patients. An approach to improve FIX replacement involves the production of more coagulatively active FIX mutants. Another promising approach in both FVIII and FIX replacement is gene therapy. Two major issues that will have to be critically addressed before gene therapy for hemophilia can become widespread are whether the procedures will be well-tolerated in patients with significant liver impairment (due to previous exposure to hepatitis viruses) and whether consistent long-term delivery of the transgenes can be achieved.

RE.CNT 35 THERE ARE 35 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L29 ANSWER 6 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 1999:677582 CAPLUS
- DN 131:306664
- TI The use of agents that by-pass factor VIII inhibitors in patients with haemophilia
- AU Roberts, Harold R.
- CS Center Thrombosis Hemostasis, School Medicine, Division Hematology Oncology, Univ. North Carolina, Chapel Hill, NC, 27599, USA
- SO Vox Sanguinis (1999), 77 (Suppl. 1), 38-41 CODEN: VOSAAD; ISSN: 0042-9007
- PB S. Karger AG
- DT Journal; General Review
- LA English
- AB A brief review with 13 refs. is given. Clin. trials of prothrombin complex concs., activated prothrombin complex concs., and recombinant factor VIIa are included. The mechanism of action of factor VIIa/tissue factor in normal coagulation reactions and of factor VIIa in the absence of factors VIII and IX is discussed. Factor VIIa alone, and in the absence of factors VIII and IX, is effective in generating thrombin on activated platelet surfaces.
- RE.CNT 13 THERE ARE 13 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

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(FILE 'HOME' ENTERED AT 11:02:09 ON 28 OCT 2004)

INDEX 'ADISCTI, ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, ANTE, AQUALINE, AQUASCI, BIOBUSINESS, BIOCOMMERCE, BIOENG, BIOSIS, BIOTECHABS, BIOTECHDS, BIOTECHNO, CABA, CANCERLIT, CAPLUS, CEABA-VTB, CEN, CIN, CONFSCI, CROPB, CROPU, DDFB, DDFU, DGENE, DISSABS, ...' ENTERED AT 11:02:32 ON 28 OCT 2004 SEA COMPOSIT? AND FACTOR IX AND FACROR VIII

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FILE 'USPATFULL' ENTERED AT 11:05:46 ON 28 OCT 2004

L1

high-purity non-immunopurified and non-nanofiltered FVIII or IX concs. than in children treated with albumin-stabilized recombinant FVIII only (OR: 22.3; CI; 7.9-62.8), independently of the other factors studied. RE.CNT 33 THERE ARE 33 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 4 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2001:751401 CAPLUS

DN 136:272923

TI Development of inhibitors in patients with **haemophilia** from India

AU Ghosh, K.; Shetty, S.; Kulkarni, B.; Nair, S.; Pawar, A.; Khare, A.;
Baindur, S.; Mohanty, D.
CS Institute of Immunohaematology, KDM Magnitude 2

CS Institute of Immunohaematology, KEM Hospital Parel, Mumbai, 400012, India SO Haemophilia (2001), 7(3), 273-278
CODEN: HAEMF4; ISSN: 1351-8216

Blackwell Science Ltd.

DT Journal

PB

LA English

- Four hundred and seven patients (352 hemophilia A and 55 hemophilia B) AB were investigated for the presence of factor VIII and IX inhibitors. Twenty-four out of 292 severe and two out of 36 moderate hemophilia A patients showed the presence of inhibitors. The mean age at development of inhibitors was 17.7 yr (range 6-52 yr). In 12 patients the inhibitors were detected due to suboptimal response to factor replacement therapy (symptomatic) and in the remaining 14 patients the inhibitors were detected during the routine screening of the patients' samples for inhibitors. They had, however, responded well to the usual doses of factor concs. and there was no suspicion in these patients that they had developed an inhibitor (asymptomatic). There were two families in which the inhibitors were detected in more than one family member. The level of inhibitors in symptomatic patients ranged from 2.2 Bethesda units (BU) mL-1 to 460.6 BU mL-1, and in asymptomatic patients it ranged from 0.8 BU mL-1 to 3.2 BU mL-1. The inhibitors persisted in all patients except one, who developed an inhibitor postoperatively for a brief period of 3 mo. All these patients were followed up from first factor exposure and were tested for inhibitors at least twice a year. The mean number of exposure days before they developed inhibitors was 47.5 exposure days (range 17-98 exposure days). No inhibitors appeared after more than 100 exposure days in any of the patients. When 50 consecutive patients were investigated for intron 22 inversions of the factor VIII gene, 17 patients were found to be pos. for inversions (10 proximal inversion; seven distal inversion) out of whom four patients developed inhibitors, three patients belonging to the same family. Out of 35 hemophilia B patients, only one patient developed an inhibitor. The overall prevalence of inhibitors was thus 8.2%, which is similar to the reports from western countries, prior to the introduction of highly purified factor concentrate therapy.
- RE.CNT 30 THERE ARE 30 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

L29 ANSWER 5 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN

AN 2000:412293 CAPLUS

DN 133:129444

TI Therapeutic approaches for haemophilia

AU Hortelano, Gonzalo; Ofosu, Frederick A.

CS Canadian Blood Services and Department of Pathology and Molecular Medicine, McMaster University, Hamilton, ON, L8N 3Z5, Can.

SO Expert Opinion on Therapeutic Patents (2000), 10(6), 929-938 CODEN: EOTPEG; ISSN: 1354-3776

PB Ashley Publications Ltd.

DT Journal; General Review

LA English

AB A review with 59 refs. The life-long episodic bleeding associated with inherited deficiencies of **blood coagulation**

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762249 FACTORS
       1358572 FACTOR
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        100449 VIII
             5 VIIIS
        100451 VIII
                 (VIII OR VIIIS)
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                 (BLOOD (W) COAGULATION (W) FACTOR (W) VIII)
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        762249 FACTORS
       1358572 FACTOR
                 (FACTOR OR FACTORS)
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             2 IXES
         71713 IX
                 (IX OR IXES)
          3376 FACTOR IX
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           425 BLOOD COAGULATION FACTOR VIII AND FACTOR IX AND PY<2003
L28
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             4 HAEMOPHILIAS
           195 HAEMOPHILIA
                  (HAEMOPHILIA OR HAEMOPHILIAS)
             6 L28 AND HAEMOPHILIA
L29
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    ANSWER 1 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
T<sub>1</sub>2.9
     2002:864083 CAPLUS
MΑ
DN
     138:71528
     Use of a non-depleting anti-CD4 antibody to modulate the immune response
     to coagulation factors VIII and IX
     Salooja, Nina; Kemball-Cook, Geoffrey; Tuddenham, Edward G. D.; Dyson,
ΑU
     Julian
     Haemostasis Research, MRC Clinical Sciences Centre, Imperial College
CS
     School of Medicine, Hammersmith Hospital, London, UK
     British Journal of Haematology (2002), 118(3), 839-842
     CODEN: BJHEAL; ISSN: 0007-1048
     Blackwell Science Ltd.
PΒ
DT
     Journal
     English
LA
     The generation of antibodies to therapeutic factors VIII or IX is a major
AB
     problem in the management of haemophilia and places potential
     limitations on the application of gene therapy. The authors have
     investigated the administration of a non-depleting anti-CD4 antibody for
     modulation of the immune response to human recombinant coagulation factors
     VIII and IX. In mice given these clotting factors, co-administration of
     anti-CD4 antibody significantly reduced the appearance of factor-specific
                  These data provide evidence that the neutralizing antibody
     response to exogenous coagulation factors may be controllable if
     non-depleting anti-CD4 antibody is co-administered at the time of initial
     replacement therapy.
               THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT 12
               ALL CITATIONS AVAILABLE IN THE RE FORMAT
L29 ANSWER 2 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
      2002:450686 CAPLUS
ΑN
      137:56965
DN
      Comparative pharmacokinetic studies in haemophilia
 TТ
      Morfini, M.
 ΑU
```

- CS Haematology Department and Haemophilia Centre, Azienda Ospedaliera Careggi, Florence, I-50134, Italy
- SO Haemophilia (2002), 8(Suppl. 2), 30-33 CODEN: HAEMF4; ISSN: 1351-8216
- PB Blackwell Science Ltd.
- DT Journal
- LA English
- The general rules of pharmacokinetics have been applied to the study of AΒ the behavior of clotting factor concs. in patients with hemophilia. Since 1980, the continuous development of innovative plasma-and rDNA-derived concs. and the implementation of new virucidal methods in manufacturing processes has prompted us to define a standard approach to this issue. Model-based methods, based upon one or two open compartment models, were available when this work was initiated. Unfortunately, these methods are supported by very little biol. data and are profoundly affected by the goodness-of-fit of the data. In contrast, the model-independent method, which is not affected by errors in fitting, provides reproducible and reliable ests. of the behavior of clotting factor concs. in patients with hemophilia. Further, the calcns. required for the model-independent method are quite simple and can be computed using a pocket minicomputer. The need for an accurate standardization has been recognized by the Factor VIII/IX Sub-Committee which, in 1991, issued the first recommendations on the pharmacokinetic evaluation of Factor VIII/IX concs. A recent revision of the recommendations has been made available on the web site of the International Society on Thrombosis and Hemostasis. The most crucial changes - sample size, study design, dosages in single-dose studies, potency assessment, need for well-defined stds., optimal number of points, and most important outcomes - are discussed in this report. In addition, the model-independent and compartmental methods are described.

RE.CNT 12 THERE ARE 12 CITED REFERENCES AVAILABLE FOR THIS RECORD ALL CITATIONS AVAILABLE IN THE RE FORMAT

- L29 ANSWER 3 OF 6 CAPLUS COPYRIGHT 2004 ACS on STN
- AN 2002:206352 CAPLUS
- DN 137:184107
- TI Prevalence of IgG antibodies to human parvovirus B19 in haemophilia children treated with recombinant factor (F)VIII only or with at least one plasma-derived FVIII or FIX concentrate: Results from the French haemophilia cohort
- AU Gaboulaud, Valerie; Parquet, Armelle; Tahiri, Cedric; Claeyssens, Segolene; Potard, Valerie; Faradji, Albert; Peynet, Jocelyne; Costagliola, Dominique
- CS Suivi Therapeutique National Des Hemophiles Group, Inserm SC4, Faculte de Medecine de Saint-Antoine, Paris, 75571/12, Fr.
- SO British Journal of Haematology (2002), 116(2), 383-389 CODEN: BJHEAL; ISSN: 0007-1048
- PB Blackwell Publishing Ltd.
- DT Journal
- LA English
- Human parvovirus B19 has been transmitted by some brands of virally AΒ attenuated plasma-derived factor VIII (FVIII) or IX (FIX) concs. To quantify the differences of human parvovirus B19 risk transmission between albumin-stabilized recombinant factor and plasma-derived factor, we studied the prevalence of IgG antibodies to B19 (anti-B19) in 193 haemophiliac children between 1 and 6-yr of age who had previously been treated with albumin-stabilized recombinant FVIII only (n = 104), and in children previously treated with solvent/detergent high-purity non-immunopurified and non-nanofiltered FVIII or IX concs. (n = 89). Association between the prevalence of anti-B19 and the treatment group was analyzed using multi-variate logistic regression. Age, severity and type of haemophilia, number of cumulative days of exposure to factor VIII or IX, previous history of red blood cells or plasma transfusion were considered as potential confounding variables. A higher prevalence of anti-B19 was found in children previously treated with solvent/detergent